

# SEARCH REQUEST FORM

Scientific and Technical Information Center

Access DB#

9/089

Requester's Full Name: Brian K. Kohn Examiner #: 78155 Date: 4/8/03  
 Art Unit: 1614 Phone Number 308-5371 Serial Number: 09/035099  
 Mail Box and Bldg/Room Location: CM1 2D04 Results Format Preferred (circle): PAPER DISK E-MAIL

If more than one search is submitted, please prioritize searches in order of need.

\*\*\*\*\*  
 Please provide a detailed statement of the search topic, and describe as specifically as possible the subject matter to be searched. Include the elected species or structures, keywords, synonyms, acronyms, and registry numbers, and combine with the concept or utility of the invention. Define any terms that may have a special meaning. Give examples or relevant citations, authors, etc, if known. Please attach a copy of the cover sheet, pertinent claims, and abstract.

Title of Invention: neurodegenerative disease

Inventors (please provide full names): Bamdad et al

Earliest Priority Filing Date: 4/12/00

\*For Sequence Searches Only\* Please include all pertinent information (parent, child, divisional, or issued patent numbers) along with the appropriate serial number.

\* method of treatment: neurodegenerative disease caused by fibril or aggregate  
 - e.g. Alzheimer, Type II diabetes, DIC (Disseminated Intravascular Coagulation), Parkinson's, Huntington's, Amyloidosis,

Jah Delaval  
 Reference Librarian  
 Biotechnology & Chemical Library  
 CM1 1E07 - 703-308-4498  
 jah.delaval@uspto.gov

## STAFF USE ONLY

Searcher: Jan  
 Searcher Phone #: 4498  
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 Date Searcher Picked Up: 4/9/03  
 Date Completed: 4/9/03  
 Searcher Prep & Review Time: \_\_\_\_\_  
 Clerical Prep Time: 30  
 Online Time: 170

## Type of Search

NA Sequence (#) \_\_\_\_\_  
 AA Sequence (#) \_\_\_\_\_  
 Structure (#) ☒  
 Bibliographic ☒  
 Litigation \_\_\_\_\_  
 Fulltext \_\_\_\_\_  
 Patent Family \_\_\_\_\_  
 Other \_\_\_\_\_

## Vendors and cost where applicable

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 Dialog \_\_\_\_\_  
 Questel/Orbit \_\_\_\_\_  
 Dr.Link \_\_\_\_\_  
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L127 ANSWER 1 OF 28 HCAPLUS COPYRIGHT 2003 ACS

AN 2002:978470 HCAPLUS

DN 138:33365

TI Compositions and methods for the treatment of **Parkinson's** disease with quinoline ring-containing **neuromelanin**-binding compounds

IN Nelson, Jodi

PA USA

SO U.S. Pat. Appl. Publ., 17 pp., Cont.-in-part of U.S. 6,417,177.

CODEN: USXXCO

DT **Patent**

LA English

IC ICM A61K031-4706

NCL 514313000

CC 1-11 (Pharmacology)

Section cross-reference(s): 63

FAN.CNT 2

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	US 2002198231	A1	20021226	US 2002-192414	20020709 <--
	US 6417177	B1	20020709	US 2000-615639	20000713 <--
PRAI	US 1999-143767P	P	19990713	<--	
	US 2000-175051P	P	20000107	<--	
	US 2000-202140P	P	20000505	<--	
	US 2000-615639	A2	20000713	<--	

AB This invention provides compns. and methods for increasing cellular respiration of melanized catecholamine **neurons**, and methods for alleviating symptoms or stopping appearance and/or progression of symptoms of **Parkinson's** disease and related conditions, characterized by nigrostriatal degeneration. An effective amt. of a **neuromelanin**-binding compn. having a quinoline ring in a suitable pharmaceutical carrier is administered to patient in need of such treatment. Preferably the compn. comprises (-)-chloroquine. Selected adjuvants are also provided as part of the compns. of this invention.

ST **Parkinson** disease treatment chloroquine compd;  
**antiparkinsonian** quinoline ring contg **neuromelanin**  
binding compd; melanized catecholamine **neuron** respiration  
chloroquine compd

IT Antihistamines  
(H1, enhancing agent adjuvant; quinoline ring-contg.  
**neuromelanin**-binding compds. for treatment of **Parkinson**  
's disease)

IT **Brain**  
(adjuvant targeting; quinoline ring-contg. **neuromelanin**  
-binding compds. for treatment of **Parkinson's** disease)

IT Antioxidants  
Dopamine agonists  
Radical scavengers  
(adjuvant; quinoline ring-contg. **neuromelanin**-binding compds.  
for treatment of **Parkinson's** disease)

IT Lactoferrins  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(antibody to, conjugates with chloroquine compd.; quinoline ring-contg.  
**neuromelanin**-binding compds. for treatment of **Parkinson**  
's disease)

IT **Nerve**  
(catecholaminergic, increasing cellular respiration of melanized;  
quinoline ring-contg. **neuromelanin**-binding compds. for  
treatment of **Parkinson's** disease)

IT Lipophilicity  
(chloroquine compd. conjugates with agent having; quinoline ring-contg.

- neuromelanin-binding compds. for treatment of Parkinson's disease)**
- IT Antibodies  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(conjugates, with chloroquine compd., to lactotransferrin; quinoline ring-contg. **neuromelanin-binding compds. for treatment of Parkinson's disease)**
- IT Respiration, animal  
(enhancement of melanized catecholamine **neurons**; quinoline ring-contg. **neuromelanin-binding compds. for treatment of Parkinson's disease)**
- IT Drug delivery systems  
(immunoconjugates, anti-lactotransferrin antibody conjugates with chloroquine compd.; quinoline ring-contg. **neuromelanin-binding compds. for treatment of Parkinson's disease)**
- IT **Nervous system**  
(multiple system atrophy; quinoline ring-contg. **neuromelanin-binding compds. for treatment of Parkinson's disease)**
- IT Melanins  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(**neuromelanins**, quinoline ring-contg. compd. binding to; quinoline ring-contg. **neuromelanin-binding compds. for treatment of Parkinson's disease)**
- IT Cytoprotective agents  
(**neuroprotectants**, adjuvant; quinoline ring-contg. **neuromelanin-binding compds. for treatment of Parkinson's disease)**
- IT **Brain, disease**  
(nigrostriatal **degeneration**; quinoline ring-contg. **neuromelanin-binding compds. for treatment of Parkinson's disease)**
- IT Metabolism, animal  
(peripheral, inhibitor of; quinoline ring-contg. **neuromelanin-binding compds. for treatment of Parkinson's disease)**
- IT Cell membrane  
(protective agent as adjuvant; quinoline ring-contg. **neuromelanin-binding compds. for treatment of Parkinson's disease)**
- IT **Antiparkinsonian agents**  
**Cognition enhancers**  
Drug delivery systems  
Enantiomers  
Human  
**Parkinson's disease**  
(quinoline ring-contg. **neuromelanin-binding compds. for treatment of Parkinson's disease)**
- IT Salts, biological studies  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(quinoline ring-contg. **neuromelanin-binding compds. for treatment of Parkinson's disease)**
- IT Mixtures  
(racemic; quinoline ring-contg. **neuromelanin-binding compds. for treatment of Parkinson's disease)**
- IT Eye  
(retina, protective agent as adjuvant; quinoline ring-contg. **neuromelanin-binding compds. for treatment of Parkinson's disease)**
- IT Drug delivery systems  
(timed-release; quinoline ring-contg. **neuromelanin-binding compds. for treatment of Parkinson's disease)**
- IT **51-61-6, Dopamine, biological studies**

- RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(adjuvant; quinoline ring-contg. **neuromelanin**-binding compds.  
for treatment of **Parkinson's** disease)
- IT 50-81-7, Vitamin C, biological studies 73-31-4, Melatonin 128-37-0,  
Butylated hydroxytoluene, biological studies 1406-18-4, Vitamin E  
9054-89-1, Superoxide dismutase 23288-49-5, Probucol 25013-16-5,  
Butylated hydroxyanisole 174882-69-0, Pycnogenol  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL  
(Biological study); USES (Uses)  
(antioxidant adjuvant; quinoline ring-contg. **neuromelanin**  
-binding compds. for treatment of **Parkinson's** disease)
- IT 299-28-5, Calcium gluconate 814-80-2, Calcium lactate 1406-16-2,  
Vitamin D 7693-13-2, Calcium citrate 10103-46-5, Calcium phosphate  
14127-61-8, Calcium ion, biological studies  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL  
(Biological study); USES (Uses)  
(as peripheral membrane protective agent; quinoline ring-contg.  
**neuromelanin**-binding compds. for treatment of **Parkinson**  
's disease)
- IT 1951-25-3, Amiodarone 51481-61-9, Cimetidine  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL  
(Biological study); USES (Uses)  
(cytochrome P 450 2D6 and A3 inhibitor inhibiting peripheral metab. of  
chloroquine compds.; quinoline ring-contg. **neuromelanin**  
-binding compds. for treatment of **Parkinson's** disease)
- IT 56-54-2, Quinidine 60-99-1, Levomepromazine 132-22-9, Chlorpheniramine  
303-49-1, Clomipramine 364-62-5, Metoclopramide 54910-89-3, Fluoxetine  
61869-08-7, Paroxetine 66357-35-5, Ranitidine 71320-77-9, Moclobemide  
79617-96-2, Sertraline 91161-71-6, Terbinafine 116644-53-2, Mibefradil  
155213-67-5, Ritonavir 169590-42-5, Celecoxib  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL  
(Biological study); USES (Uses)  
(cytochrome P 450 2D6 inhibitor inhibiting peripheral metab. of  
chloroquine compds.; quinoline ring-contg. **neuromelanin**  
-binding compds. for treatment of **Parkinson's** disease)
- IT 114-07-8, Erythromycin 147-84-2, biological studies 42399-41-7,  
Diltiazem 54739-18-3, Fluvoxamine 65277-42-1, Ketoconazole  
70458-96-7, Norfloxacin 81103-11-9, Clarithromycin 83366-66-9,  
Nefazodone 83891-03-6, Norfluoxetine 84371-65-3, Mifepristone  
84625-61-6, Itraconazole 85721-33-1, Ciprofloxacin 86386-73-4,  
Fluconazole 127779-20-8, Saquinavir 136817-59-9, Delavirdine  
150378-17-9, Indinavir 159989-64-7, Nelfinavir  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL  
(Biological study); USES (Uses)  
(cytochrome P 450 A3 inhibitor inhibiting peripheral metab. of  
chloroquine compds.; quinoline ring-contg. **neuromelanin**  
-binding compds. for treatment of **Parkinson's** disease)
- IT 58-33-3, Promethazine hydrochloride 58-73-1, Diphenhydramine 59-33-6,  
Pyrilamine maleate 91-81-6, Tripelenamine 113-92-8, Chlorpheniramine  
maleate 303-25-3, Cyclizine hydrochloride 523-87-5, Dimenhydrinate  
980-71-2, Brompheniramine maleate 1104-22-9, Meclizine hydrochloride  
2192-20-3, Hydroxyzine hydrochloride 3505-38-2, Carbinoxamine maleate  
5897-19-8, Cyclizine lactate 10246-75-0, Hydroxyzine pamoate  
15686-51-8, Clemastine 50679-08-8, Terfenadine 68844-77-9, Astemizole  
79794-75-5, Loratadine 83881-52-1, Cetirizine hydrochloride  
87848-99-5, Acrivastine  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL  
(Biological study); USES (Uses)  
(**histamine** H1 receptor antagonist as enhancing agent  
adjuvant; quinoline ring-contg. **neuromelanin**-binding compds.  
for treatment of **Parkinson's** disease)
- IT 329322-82-9, Cytochrome P450 3A 330597-62-1, Cytochrome P450 2D6  
RL: BSU (Biological study, unclassified); BIOL (Biological study)



(inhibitor of; quinoline ring-contg. **neuromelanin**-binding compds. for treatment of **Parkinson's** disease)

IT 50-63-5, Chloroquine phosphate 54-05-7, Chloroquine 118-42-3, Hydroxychloroquine 134-31-6, 8-Quinololinol sulfate 442-96-6 1915-92-0 2739-16-4 4169-19-1, 1-Acetyl-1,2,3,4-tetrahydroquinoline 4298-15-1 6168-85-0 24283-71-4, 1-Butyryl-1,2,3,4-tetrahydroquinoline 32571-37-2 53462-15-0 58175-87-4, (-)-Chloroquine 82351-01-7 99218-67-4 319912-96-4 319912-97-5 319912-98-6 319912-99-7 319913-00-3 319913-01-4 319913-03-6 319913-04-7 319913-05-8 319913-06-9 319913-07-0 319913-08-1 478784-57-5 478784-58-6 478784-60-0 478784-61-1 478784-63-3 478784-64-4 478784-65-5 478784-66-6 478784-67-7 478784-68-8 478784-70-2 478784-71-3 478784-73-5 478784-74-6

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

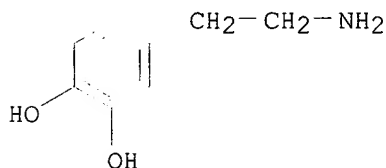
(quinoline ring-contg. **neuromelanin**-binding compds. for treatment of **Parkinson's** disease)

IT 51-61-6, Dopamine, biological studies

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (adjuvant; quinoline ring-contg. **neuromelanin**-binding compds. for treatment of **Parkinson's** disease)

RN 51-61-6 HCAPLUS

CN 1,2-Benzenediol, 4-(2-aminoethyl)- (9CI) (CA INDEX NAME)

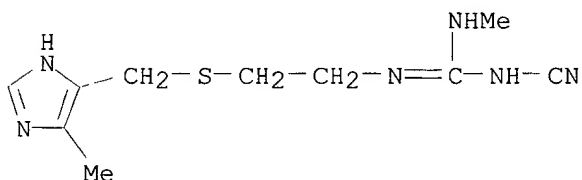


IT 51481-61-9, Cimetidine

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (cytochrome P 450 2D6 and A3 inhibitor inhibiting peripheral metab. of chloroquine compds.; quinoline ring-contg. **neuromelanin**-binding compds. for treatment of **Parkinson's** disease)

RN 51481-61-9 HCAPLUS

CN Guanidine, N-cyano-N'-methyl-N''-[2-[[5-methyl-1H-imidazol-4-yl)methyl]thio]ethyl]- (9CI) (CA INDEX NAME)



L127 ANSWER 2 OF 28 HCAPLUS COPYRIGHT 2003 ACS

AN 2002:793363 HCAPLUS

DN 137:304808

TI Methods and compounds for decreasing **cell toxicity** or death

IN Yuan, Junying; Sanchez, Ivelisse

PA President and Fellows of Harvard College, USA

SO PCT Int. Appl., 137 pp.

CODEN: PIXXD2

DT Patent  
 LA English  
 IC ICM A61K  
 CC 1-12 (Pharmacology)  
 FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2002080855	A2	20021017	WO 2002-US11025	20020409
	W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
	US 2002155172	A1	20021024	US 2001-829040	20010409 <--
PRAI	US 2001-829040	A	20010409		
	US 2000-195661P	P	20000407	<--	
AB	The invention features methods for decreasing cell toxicity or death, and for decreasing polyglutamine <b>aggregates</b> and other <b>amyloidogenic aggregates</b> . The invention also features methods for treating a subject with a condition in which expanded polyglutamine repeats or <b>amyloidogenic aggregates</b> are present.				
ST	cell death inhibitor polyglutamine <b>aggregate</b> decrease; <b>amyloidogenic aggregate</b> decrease cell death inhibitor; Congo Red analog cell death polyglutamine <b>aggregate</b>				
IT	<b>Nervous system</b> (Huntington's chorea; methods and compds. for decreasing cell toxicity or death by decreasing polyglutamine or <b>amyloidogenic</b> proteins <b>aggregates</b> using Congo Red and related compds. in relation to mechanism)				
IT	<b>Proteins</b> RL: BSU (Biological study, unclassified); BIOL (Biological study) ( <b>amyloidogenic</b> ; methods and compds. for decreasing cell toxicity or death by decreasing polyglutamine or <b>amyloidogenic</b> proteins <b>aggregates</b> using Congo Red and related compds. in relation to mechanism)				
IT	<b>Human</b> Mammalia Rodentia (cell; methods and compds. for decreasing cell toxicity or death by decreasing polyglutamine or <b>amyloidogenic</b> proteins <b>aggregates</b> using Congo Red and related compds. in relation to mechanism)				
IT	<b>Nervous system</b> (degeneration; methods and compds. for decreasing cell toxicity or death by decreasing polyglutamine or <b>amyloidogenic</b> proteins <b>aggregates</b> using Congo Red and related compds. in relation to mechanism)				
IT	<b>Brain, disease</b> (dentatorubral-pallidoluysian atrophy; methods and compds. for decreasing cell toxicity or death by decreasing polyglutamine or <b>amyloidogenic</b> proteins <b>aggregates</b> using Congo Red and related compds. in relation to mechanism)				
IT	<b>Schizophrenia</b> (familial; methods and compds. for decreasing cell toxicity or death by decreasing polyglutamine or <b>amyloidogenic</b> proteins <b>aggregates</b> using Congo Red and related compds. in relation to mechanism)				
IT	<b>Fertility</b>				

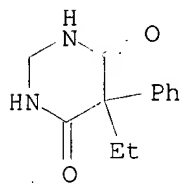
- (male, disorder; methods and compds. for decreasing cell toxicity or death by decreasing polyglutamine or **amyloidogenic** proteins **aggregates** using Congo Red and related compds. in relation to mechanism)
- IT Cell death  
Clover (*Trifolium pratense*)  
Cytoprotective agents  
Cytotoxicity  
Drug delivery systems  
Drug screening  
Gamete and Germ cell  
High throughput screening  
Translation, genetic  
(methods and compds. for decreasing cell toxicity or death by decreasing polyglutamine or **amyloidogenic** proteins **aggregates** using Congo Red and related compds. in relation to mechanism)
- IT Spinal muscular atrophy  
(spinal and bulbar muscular atrophy; methods and compds. for decreasing cell toxicity or death by decreasing polyglutamine or **amyloidogenic** proteins **aggregates** using Congo Red and related compds. in relation to mechanism)
- IT Nervous system  
(spinocerebellar ataxia; methods and compds. for decreasing cell toxicity or death by decreasing polyglutamine or **amyloidogenic** proteins **aggregates** using Congo Red and related compds. in relation to mechanism)
- IT 2829-42-7, Direct Yellow 26  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(Direct Yellow 26; methods and compds. for decreasing cell toxicity or death by decreasing polyglutamine or **amyloidogenic** proteins **aggregates** using Congo Red and related compds. in relation to mechanism)
- IT 179241-78-2, Caspase 8  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(methods and compds. for decreasing cell toxicity or death by decreasing polyglutamine or **amyloidogenic** proteins **aggregates** using Congo Red and related compds. in relation to mechanism)
- IT 573-58-0, Congo Red  
RL: DMA (Drug mechanism of action); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(methods and compds. for decreasing cell toxicity or death by decreasing polyglutamine or **amyloidogenic** proteins **aggregates** using Congo Red and related compds. in relation to mechanism)
- IT 50-23-7, Hydrocortisone 52-86-8, Haloperidol 59-99-4D, Neostigmine, derivs. 68-41-7, D-Cycloserine 92-87-5D, [1,1'-Biphenyl]-4,4'-diamine, derivs. 94-78-0, Phenazopyridine 125-33-7, Primidone 446-86-6, Azathioprine 613-35-4 1309-48-4, Magnesium oxide, biological studies 2429-84-7, Direct Red 1 2444-46-4, N-Vanillylnonamide 2623-51-0 2921-57-5 6459-87-6 6637-88-3, Direct Orange 6 6655-95-4, Direct Blue 158 8005-72-9, Direct Yellow 28 14611-52-0, (-)-Deprenyl hydrochloride 20830-75-5, Digoxin 22260-51-1, Bromocriptine mesylate 34977-63-4, Direct Black 51 42924-53-8, Nabumetone 54579-28-1, C.I. Direct Orange 1 64083-59-6, Direct Orange 8 75535-02-3 78374-79-5 82859-73-2 109019-05-8 109504-77-0 299199-67-0 303215-90-9 305858-09-7 312749-40-9 330220-26-3 331856-01-0 349653-80-1 351987-68-3 364600-08-8 385790-72-7 404852-48-8 413617-33-1 414887-35-7 470688-32-5 470688-33-6 470688-34-7 470688-35-8D, derivs. 470688-36-9D, derivs. 470688-37-0 470697-02-0 470697-03-1 470697-04-2

RL: PAC (Pharmacological activity); **THU (Therapeutic use)**; BIOL (Biological study); USES (Uses)  
 (methods and compds. for decreasing cell toxicity or death by decreasing polyglutamine or **amyloidogenic** proteins **aggregates** using Congo Red and related compds. in relation to mechanism)

IT 26700-71-0, Polyglutamine 69864-43-3, Polyglutamine  
 RL: BSU (Biological study, unclassified); BIOL (Biological study)  
 (repeats; methods and compds. for decreasing cell toxicity or death by decreasing polyglutamine or **amyloidogenic** proteins **aggregates** using Congo Red and related compds. in relation to mechanism)

IT 125-33-7, Primidone  
 RL: PAC (Pharmacological activity); **THU (Therapeutic use)**; BIOL (Biological study); USES (Uses)  
 (methods and compds. for decreasing cell toxicity or death by decreasing polyglutamine or **amyloidogenic** proteins **aggregates** using Congo Red and related compds. in relation to mechanism)

RN 125-33-7 HCAPLUS  
 CN 4,6(1H,5H)-Pyrimidinedione, 5-ethylhydro-5-phenyl- (6CI, 8CI, 9CI) (CA INDEX NAME)



L127 ANSWER 3 OF 28 HCAPLUS COPYRIGHT 2003 ACS  
 AN 2002:522639 HCAPLUS  
 DN 137:73276  
 TI Behavior chemotherapy for prevention of **Alzheimer's** disease  
 IN Eig, Mark H.  
 PA USA  
 SO U.S. Pat. Appl. Publ., 8 pp., Cont.-in-part of U.S. 6,333,357.  
 CODEN: USXXCO

DT **Patent**  
 LA English  
 IC ICM A61K031-55  
 ICS A61K031-44; A61K031-343; A61K031-137  
 NCL 514355000  
 CC 1-11 (Pharmacology)  
 FAN.CNT 2

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	US 2002091145	A1	20020711	US 2001-992972	20011113 <--
	US 6333357	B1	20011225	US 1999-434286	19991105 <--
	AU 2001018105	A5	20020611	AU 2001-18105	20001201 <--
PRAI	US 1999-434286	A2	19991105	<--	
	WO 2000-US32697	A	20001201	<--	

AB A protocol for prevention of **Alzheimer's** disease onset is described. The protocol involves stimulating the implicit memory, followed by continuing such stimulation in conjunction with psychol. treatments followed by continuing the stimulation of the implicit memory and, in addn., stimulating the explicit memory. Use of this protocol results in a permanent replacement of undesirable behaviors with desirable ones.

ST behavior chemotherapy **Alzheimer** disease memory stimulation

IT **Alzheimer's disease**  
**Anti-Alzheimer's agents**  
 Antihypertensives  
 Behavior  
 Cardiovascular agents  
**Cognition enhancers**  
 Human  
 Hypertension  
 Memory, biological  
 Psychotropics  
 Vasodilators  
 (behavior chemotherapy for prevention of **Alzheimer's** disease)

IT Corticosteroids, biological studies  
 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (behavior chemotherapy for prevention of **Alzheimer's** disease)

IT Rhythm, biological  
 (circadian, diurnal metab.; behavior chemotherapy for prevention of **Alzheimer's** disease)

IT Metabolism, animal  
 (diurnal; behavior chemotherapy for prevention of **Alzheimer's** disease)

IT Medicine  
 (psychol., psychol. treatment; behavior chemotherapy for prevention of **Alzheimer's** disease)

IT Heart, disease  
 (tachycardia; behavior chemotherapy for prevention of **Alzheimer's** disease)

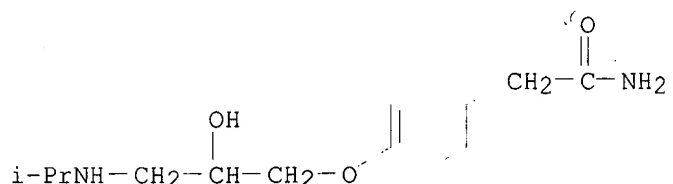
IT Adrenoceptor antagonists  
 (.beta.-; behavior chemotherapy for prevention of **Alzheimer's** disease)

IT 122-09-8, Phentermine 458-24-2, Fenfluramine 1953-04-4, Galantamine hydrobromide 21829-25-4, Nifedipine **29122-68-7**, Atenolol 59729-33-8, Citalopram 111470-99-6, Amlodipine besylate 120014-06-4, Donepezil 129101-54-8  
 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (behavior chemotherapy for prevention of **Alzheimer's** disease)

IT **29122-68-7**, Atenolol  
 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (behavior chemotherapy for prevention of **Alzheimer's** disease)

RN **29122-68-7** HCAPLUS

CN Benzeneacetamide, 4-[2-hydroxy-3-[(1-methylethyl)amino]propoxy]- (9CI)  
 (CA INDEX NAME)



L127 ANSWER 4 OF 28 HCAPLUS COPYRIGHT 2003 ACS

AN 2002:429537 HCAPLUS

DN 137:748

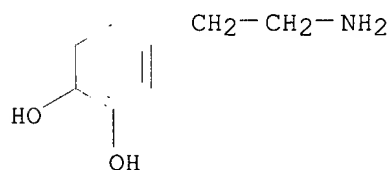
TI Method using melatonin inhibitors for the treatment of **neurological** or **neuropsychiatric** disorders

IN Willis, Gregory Lynn  
 PA Australia  
 SO U.S. Pat. Appl. Publ., 34 pp., Cont.-in-part of U.S. Ser. No. 285,859.  
 CODEN: USXXCO  
 DT **Patent**  
 LA English  
 IC ICM A61K031-00  
 NCL 514001000  
 CC 1-11 (Pharmacology)  
 FAN.CNT 4

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	US 2002068692	A1	20020606	US 2001-971783	20011009 <--
	WO 9815267	A1	19980416	WO 1997-AU661	19971003 <--
	W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
	RW: GH, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG				
	US 6310085	B1	20011030	US 1999-285859	19990402 <--
PRAI	AU 1996-2745	A	19961004 <--		
	WO 1997-AU661	A2	19971003 <--		
	US 1999-285859	A2	19990402 <--		
AB	A method for the treatment and/or prophylaxis of a <b>neurol.</b> or <b>neuropsychiatric</b> disorder assocd. with altered dopamine function is disclosed which comprises subjecting a patient in need thereof to therapy which blocks and/or inhibits melatonin, precursors thereof, and/or metabolic products thereof.				
ST	<b>neurol neuropsychiatric</b> disorder treatment melatonin inhibitor; dopamine <b>neurol</b> disorder treatment melatonin inhibitor				
IT	<b>Brain, disease</b> (Gilles de la Tourette syndrome; melatonin inhibitors for treatment of <b>neurol.</b> or <b>neuropsychiatric</b> disorder)				
IT	<b>Nervous system</b> (Huntington's chorea; melatonin inhibitors for treatment of <b>neurol.</b> or <b>neuropsychiatric</b> disorder)				
IT	<b>Mental disorder</b> (Pick's disease; melatonin inhibitors for treatment of <b>neurol.</b> or <b>neuropsychiatric</b> disorder)				
IT	<b>Disease, animal</b> (Sundowner's syndrome; melatonin inhibitors for treatment of <b>neurol.</b> or <b>neuropsychiatric</b> disorder)				
IT	<b>Stress, animal</b> (acute stress disorder; melatonin inhibitors for treatment of <b>neurol.</b> or <b>neuropsychiatric</b> disorder)				
IT	<b>Mental disorder</b> (agoraphobia; melatonin inhibitors for treatment of <b>neurol.</b> or <b>neuropsychiatric</b> disorder)				
IT	<b>Nervous system</b> (akathisia; melatonin inhibitors for treatment of <b>neurol.</b> or <b>neuropsychiatric</b> disorder)				
IT	<b>Anorexia</b> (anorexia cachexia; melatonin inhibitors for treatment of <b>neurol.</b> or <b>neuropsychiatric</b> disorder)				
IT	<b>Appetite</b> (anorexia <b>nervosa</b> ; melatonin inhibitors for treatment of <b>neurol.</b> or <b>neuropsychiatric</b> disorder)				
IT	<b>Ion channel blockers</b>				

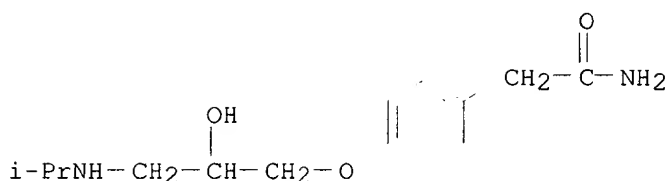
- (calcium; melatonin inhibitors for treatment of **neurol. or neuropsychiatric disorder**)
- IT **Mental disorder**
  - (**dementia**; melatonin inhibitors for treatment of **neurol. or neuropsychiatric disorder**)
- IT **Nervous system**
  - (disease; melatonin inhibitors for treatment of **neurol. or neuropsychiatric disorder**)
- IT Toxicity
  - (drug, **neuroleptic-induced parkinsonism**; melatonin inhibitors for treatment of **neurol. or neuropsychiatric disorder**)
- IT **Nervous system**
  - (**dystonia**, acute; melatonin inhibitors for treatment of **neurol. or neuropsychiatric disorder**)
- IT Anxiety
  - (generalized anxiety disorder; melatonin inhibitors for treatment of **neurol. or neuropsychiatric disorder**)
- IT Disease, animal
  - (malignant syndrome; melatonin inhibitors for treatment of **neurol. or neuropsychiatric disorder**)
- IT **Anti-Alzheimer's agents**
  - Antidepressants
  - Antiparkinsonian agents**
  - Antipsychotics
  - Anxiolytics
  - Diagnosis
  - Mental disorder**
  - Movement disorders
  - Nervous system agents**
  - Phototherapy
  - Psychotropics
  - Schizophrenia**
  - Wernicke-Korsakoff syndrome
    - (melatonin inhibitors for treatment of **neurol. or neuropsychiatric disorder**)
- IT Melatonin receptors
  - RL: BSU (Biological study, unclassified); BIOL (Biological study)
  - (melatonin inhibitors for treatment of **neurol. or neuropsychiatric disorder**)
- IT Tranquilizers
  - (**neuroleptic-induced parkinsonism**; melatonin inhibitors for treatment of **neurol. or neuropsychiatric disorder**)
- IT **Mental disorder**
  - (obsession-compulsion; melatonin inhibitors for treatment of **neurol. or neuropsychiatric disorder**)
- IT Anxiety
  - (panic; melatonin inhibitors for treatment of **neurol. or neuropsychiatric disorder**)
- IT Leg
  - (periodic limb movement syndrome and restless leg syndrome; melatonin inhibitors for treatment of **neurol. or neuropsychiatric disorder**)
- IT Arm
  - (periodic limb movement syndrome; melatonin inhibitors for treatment of **neurol. or neuropsychiatric disorder**)
- IT **Mental disorder**
  - (post-traumatic stress disorder; melatonin inhibitors for treatment of **neurol. or neuropsychiatric disorder**)
- IT Paralysis
  - (progressive subnuclear palsy; melatonin inhibitors for treatment of **neurol. or neuropsychiatric disorder**)

- IT Disease, animal  
(punch drunk syndrome; melatonin inhibitors for treatment of **neurol. or neuropsychiatric disorder**)
- IT Brain, disease  
(**stroke**; melatonin inhibitors for treatment of **neurol. or neuropsychiatric disorder**)
- IT Pineal gland  
(surgical ablation or destruction; melatonin inhibitors for treatment of **neurol. or neuropsychiatric disorder**)
- IT Nervous system  
(tardive dyskinesia; melatonin inhibitors for treatment of **neurol. or neuropsychiatric disorder**)
- IT Multiple sclerosis  
(therapeutic agents; melatonin inhibitors for treatment of **neurol. or neuropsychiatric disorder**)
- IT Anti-ischemic agents  
(trans-ischemic attack; melatonin inhibitors for treatment of **neurol. or neuropsychiatric disorder**)
- IT Adrenoceptor antagonists  
(.beta.-; melatonin inhibitors for treatment of **neurol. or neuropsychiatric disorder**)
- IT 73-31-4, Melatonin  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(and precursors and metabolites; melatonin inhibitors for treatment of **neurol. or neuropsychiatric disorder**)
- IT 51-61-6, Dopamine, biological studies  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(melatonin inhibitors for treatment of **neurol. or neuropsychiatric disorder**)
- IT 525-66-6, Propranolol 9002-79-3, Melanocyte-stimulating hormone  
29122-68-7, Atenolol 115007-18-6, ML-23  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL  
(Biological study); USES (Uses)  
(melatonin inhibitors for treatment of **neurol. or neuropsychiatric disorder**)
- IT 51-61-6, Dopamine, biological studies  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(melatonin inhibitors for treatment of **neurol. or neuropsychiatric disorder**)
- RN 51-61-6 HCAPLUS
- CN 1,2-Benzenediol, 4-(2-aminoethyl)- (9CI) (CA INDEX NAME)



- IT 29122-68-7, Atenolol  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL  
(Biological study); USES (Uses)  
(melatonin inhibitors for treatment of **neurol. or neuropsychiatric disorder**)
- RN 29122-68-7 HCAPLUS
- CN Benzeneacetamide, 4-[2-hydroxy-3-[(1-methylethyl)amino]propoxy]- (9CI)  
(CA INDEX NAME)





L127 ANSWER 5 OF 28 HCAPLUS COPYRIGHT 2003 ACS

AN 2002:368310 HCAPLUS

DN 136:363866

TI Serotonergic compositions and methods for treatment of mild **cognitive** impairment

IN Wurtman, Richard J.; Lee, Robert K. K.

PA Massachusetts Institute of Technology, USA

SO PCT Int. Appl., 34 pp.

CODEN: PIXXD2

DT **Patent**

LA English

IC ICM A61K031-00

CC 1-11 (Pharmacology)

Section cross-reference(s): 63

FAN.CNT 2

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE	
PI	WO 2002038142	A2	20020516	WO 2001-US43016	20011108	<--
	W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
	RW:	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
	AU 2002030423	A5	20020521	AU 2002-30423	20011108	<--
	US 2002173511	A1	20021121	US 2001-986469	20011108	<--
	US 2002173549	A1	20021121	US 2001-986470	20011108	<--
PRAI	US 2000-246615P	P	20001108			<--
	WO 2001-US43016	W	20011108			
AB	A method of treating mild <b>cognitive</b> impairment is disclosed. The method comprises administering an effective amt. of a serotonergic agent, including, but not limited to, dextnorfenfluramine. The agent can be any serotonergic agonist, partial agonist, serotonin reuptake inhibitor, or combinations of these agents. The treatment method also encompasses combinations of serotonergic agents and nonsteroidal antiinflammatory agents. The treatment method may also delay the onset of mild <b>cognitive</b> impairment, <b>dementia</b> , or both.					
ST	mild <b>cognitive</b> impairment <b>dementia</b> serotonergic agent pharmaceutical; serotonin reuptake inhibitor mild <b>cognitive</b> impairment; dextnorfenfluramine mild <b>cognitive</b> impairment; serotonergic agent NSAID mild <b>cognitive</b> impairment					
IT	<b>Astrocyte</b> (APP overexpression inhibition in; serotonergic compns. and methods for treatment of mild <b>cognitive</b> impairment)					
IT	5-HT agonists (and partial agonists; serotonergic compns. and methods for treatment of mild <b>cognitive</b> impairment)					
IT	Drug delivery systems (enteric; serotonergic compns. and methods for treatment of mild					

**cognitive** impairment)

IT Drug delivery systems  
(oral; serotonergic compns. and methods for treatment of mild **cognitive** impairment)

IT Drug delivery systems  
(parenterals; serotonergic compns. and methods for treatment of mild **cognitive** impairment)

IT 5-HT antagonists  
**Cognition enhancers**  
Drug delivery systems  
(serotonergic compns. and methods for treatment of mild **cognitive** impairment)

IT Cerebrospinal fluid  
(sol. **amyloid** precursor protein in; serotonergic compns. and methods for treatment of mild **cognitive** impairment)

IT **Amyloid precursor proteins**  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(sol., secretion-stimulating agent; serotonergic compns. and methods for treatment of mild **cognitive** impairment)

IT Drug delivery systems  
(topical; serotonergic compns. and methods for treatment of mild **cognitive** impairment)

IT 50-67-9, Serotonin, biological studies  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(reuptake inhibitors; serotonergic compns. and methods for treatment of mild **cognitive** impairment)

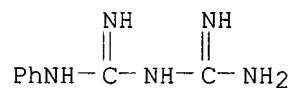
IT 66-83-1 73-22-3, Tryptophan, biological studies 93-65-2, MCPP  
102-02-3, Phenylbiguanide 153-98-0, Serotonin hydrochloride  
303-49-1, Clomipramine 304-52-9, .alpha.-Methyl-5-hydroxytryptamine  
458-24-2, Fenfluramine 971-74-4, Serotonin creatinine sulfate  
1054-88-2, Spiroxatrine 1152-76-7, Mescaline sulfate 1814-64-8,  
LY-165163 2113-05-5 2170-58-3 2315-02-8, Oxymetazoline hydrochloride  
2963-79-3, Bufotenine monooxalate 3036-16-6, Serotonin oxalate  
3239-44-9, Dexfenfluramine 4350-09-8, 5-Hydroxy tryptophan 4774-24-7,  
Quipazine 5464-78-8, 1-(2-Methoxyphenyl)piperazine hydrochloride  
5787-02-0 13078-15-4, 1-(3-Chlorophenyl)piperazine hydrochloride  
15232-63-0 15532-75-9, TFMPP 17286-40-7 19036-73-8,  
Dexnorfenfluramine 21102-95-4, BMY 7378 24973-25-9 28614-26-8,  
N-Methylquipazine 29705-96-2, **34661-85-3**, 5-Methylurapidil  
36505-84-7, Buspirone 42203-78-1 48144-44-1, m-Chlorophenylbiguanide  
54739-18-3, Fluvoxamine 54910-89-3, Fluoxetine 57477-39-1  
59729-33-8, Citalopram 61869-08-7, Paroxetine 64022-27-1, MK 212  
64887-14-5, Urapidil hydrochloride 74885-09-9, 5-Carboxamidotryptamine  
74885-25-9 76135-31-4 77145-61-0 77372-73-7, 6-Nitroquipazine  
78095-20-2 78263-90-8, 2-Methyl-5-hydroxytryptamine 78263-91-9  
78950-78-4 79617-96-2, Sertraline 80300-09-0 82864-02-6  
82900-57-0, BP-554 98330-05-3, Anpirtoline 99665-05-1 103628-46-2,  
Sumatriptan 107008-28-6, RU 24969 **109028-10-6**, CGS-12066B  
109140-25-2 124756-23-6, MDL 73005EF 127126-20-9 127792-75-0, CP  
93129 134296-40-5 137403-12-4 141196-99-8, SC 53116 141510-98-7  
143137-35-3, RS 56812 148868-55-7 150323-78-7, Quipazine dimaleate  
155106-73-3, 2-[1-(4-Piperonyl)piperazinyl]benzothiazole 157798-12-4,  
5-Nonyloxytryptamine 157798-13-5 159559-70-3 160521-72-2  
160845-95-4 168986-60-5, RS 67333 169675-08-5 171205-17-7  
182563-09-3 185259-85-2, GR 46611 187665-60-7  
RL: PAC (Pharmacological activity); **THU (Therapeutic use)**; BIOL  
(Biological study); USES (Uses)  
(serotonergic compns. and methods for treatment of mild **cognitive** impairment)

IT **102-02-3**, Phenylbiguanide **34661-85-3**, 5-Methylurapidil  
**109028-10-6**, CGS-12066B  
RL: PAC (Pharmacological activity); **THU (Therapeutic use)**; BIOL  
(Biological study); USES (Uses)

(serotonergic compns. and methods for treatment of mild  
cognitive impairment)

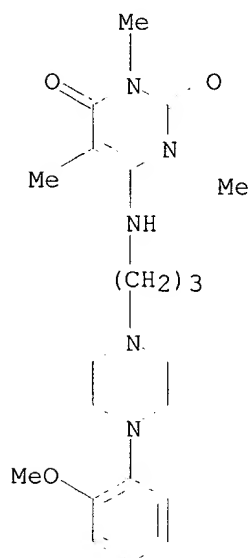
RN 102-02-3 HCAPLUS

CN Imidodicarbonimidic diamide, N-phenyl- (9CI) (CA INDEX NAME)



RN 34661-85-3 HCAPLUS

CN 2,4(1H,3H)-Pyrimidinedione, 6-[[3-[4-(2-methoxyphenyl)-1-piperazinyl]propyl]amino]-1,3,5-trimethyl- (9CI) (CA INDEX NAME)



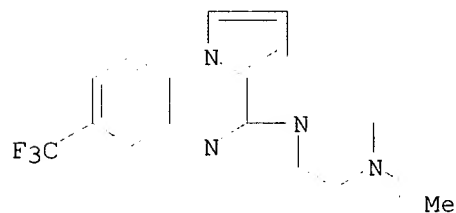
RN 109028-10-6 HCAPLUS

CN Pyrrolo[1,2-a]quinoxaline, 4-(4-methyl-1-piperazinyl)-7-(trifluoromethyl)-, (2Z)-2-butenedioate (1:2) (9CI) (CA INDEX NAME)

CM 1

CRN 109028-09-3

CMF C17 H17 F3 N4

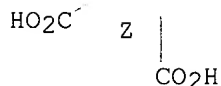


CM 2

CRN 110-16-7

CMF C4 H4 O4

Double bond geometry as shown.



L127 ANSWER 6 OF 28 HCAPLUS COPYRIGHT 2003 ACS

AN 2002:314758 HCAPLUS

DN 136:319416

TI Combination of acetylcholinesterase inhibitors and GABAA inverse agonists for the treatment of **cognitive** disorders

IN Villalobos, Anabella; Cassella, James Vincent; Rajachandran, Lavanya

PA Pfizer Products Inc., USA; Neurogen Corporation

SO PCT Int. Appl., 32 pp.

CODEN: PIXXD2

DT **Patent**

LA English

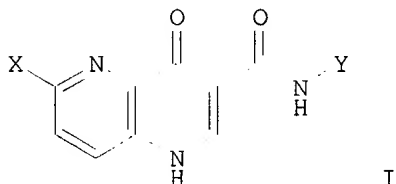
IC ICM A61K031-00

CC 1-11 (Pharmacology)

Section cross-reference(s): 63

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE	
PI	WO 2002032412	A2	20020425	WO 2001-IB1934	20011015	<--
	W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
	RW:	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
	US 2002151591	A1	20021017	US 2001-976347	20011012	<--
	AU 2001094117	A5	20020429	AU 2001-94117	20011015	<--
PRAI	US 2000-241145P	P	20001017	<--		
	WO 2001-IB1934	W	20011015			
OS	MARPAT 136:319416					
GI						



AB This invention provides a compn. for treating a **cognitive** disorder, which comprises an acetylcholinesterase, and a GABAA inverse agonist selected from a compd. (I, where X = e.g., H, halo, Ph, naphthyl, pyridinyl; Y = e.g., C1-8 alkyl, carbocycle). Thus, aricept and a GABAA inverse agonist (e.g., N-benzyl-6-ethoxy-4-oxo-1,4-tetrahydro-1,5-naphthyridine-3-carboxamide), when coadministered, interact to attenuate scopolamine-induced deficits in the spatial water maze.

ST acetylcholinesterase inhibitor GABAA agonist **cognitive** enhancer;

aricept tetrahydronaphthyridinecarboxamide **cognitive** enhancer

IT GABA agonists  
(GABAA; combination of acetylcholinesterase inhibitors and GABAA inverse agonists for treatment of **cognitive** disorders)

IT **Mental disorder**  
(attention deficit disorder; combination of acetylcholinesterase inhibitors and GABAA inverse agonists for treatment of **cognitive** disorders)

IT **Mental disorder**  
(**cognitive**; combination of acetylcholinesterase inhibitors and GABAA inverse agonists for treatment of **cognitive** disorders)

IT Anti-Alzheimer's agents  
Antiparkinsonian agents  
Cognition enhancers  
Down's syndrome  
Drug delivery systems  
Memory, biological  
Psychotropics  
(combination of acetylcholinesterase inhibitors and GABAA inverse agonists for treatment of **cognitive** disorders)

IT Mental disorder  
(~~dementia~~, vascular; combination of acetylcholinesterase inhibitors and GABAA inverse agonists for treatment of **cognitive** disorders)

IT Cognition  
(disorder; combination of acetylcholinesterase inhibitors and GABAA inverse agonists for treatment of **cognitive** disorders)

IT Drug delivery systems  
(prodrugs; combination of acetylcholinesterase inhibitors and GABAA inverse agonists for treatment of **cognitive** disorders)

IT Brain, disease  
(**stroke**; combination of acetylcholinesterase inhibitors and GABAA inverse agonists for treatment of **cognitive** disorders)

IT Drug interactions  
(synergistic; combination of acetylcholinesterase inhibitors and GABAA inverse agonists for treatment of **cognitive** disorders)

IT **Brain, disease**  
(trauma; combination of acetylcholinesterase inhibitors and GABAA inverse agonists for treatment of **cognitive** disorders)

IT 52-68-6, Metrifonate 57-47-6, Physostigmine 321-64-2, Tacrine  
357-70-0, Galantamine 102518-79-6, Huperzine A 120011-70-3, Aricept  
123441-03-2, Rivastigmine 145508-78-7, Icopezil 220860-40-2  
220860-45-7 220860-47-9 220860-48-0 220860-50-4 220860-52-6  
220860-56-0 220860-58-2 220860-59-3 220860-60-6 220860-62-8  
220860-64-0 220860-65-1 220860-66-2 220860-67-3 220860-68-4  
220860-70-8 220860-72-0 220860-74-2 220860-75-3 220860-81-1  
220860-90-2 220860-92-4 220860-93-5 220860-95-7 220861-17-6  
304680-79-3 415678-14-7 415678-15-8  
RL: PAC (Pharmacological activity); THU (**Therapeutic use**); BIOL  
(Biological study); USES (Uses)  
(combination of acetylcholinesterase inhibitors and GABAA inverse agonists for treatment of **cognitive** disorders)

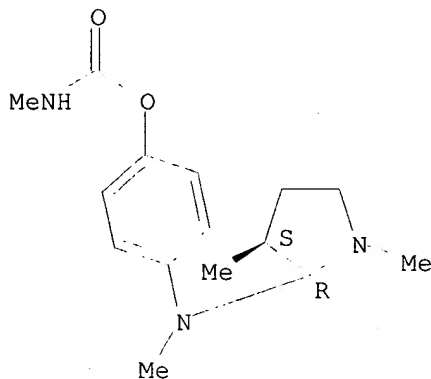
IT 9000-81-1, Acetylcholinesterase  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(inhibitors; combination of acetylcholinesterase inhibitors and GABAA inverse agonists for treatment of **cognitive** disorders)

IT **57-47-6, Physostigmine**  
RL: PAC (Pharmacological activity); THU (**Therapeutic use**); BIOL  
(Biological study); USES (Uses)  
(combination of acetylcholinesterase inhibitors and GABAA inverse agonists for treatment of **cognitive** disorders)

RN 57-47-6 HCAPLUS

CN Pyrrolo[2,3-b]indol-5-ol, 1,2,3,3a,8,8a-hexahydro-1,3a,8-trimethyl-, methylcarbamate (ester), (3aS,8aR)- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).



L127 ANSWER 7 OF 28 HCAPLUS COPYRIGHT 2003 ACS

AN 2002:241334 HCAPLUS

DN 136:257275

TI Method and composition for modulating amyloidosis

IN Reiner, Peter B.; Lam, Fred Chiu-Lai

PA Can.

SO U.S. Pat. Appl. Publ., 38 pp., Cont.-in-part of U.S. Ser. No. 67,523, abandoned.

CODEN: USXXCO

DT Patent

LA English

IC ICM C12Q001-00

ICS G01N033-53; A61K038-00; G01N033-00; C12N009-00; C07K005-00; C07K007-00; C07K016-00; C07K017-00; A61K038-12

NCL 514011000

CC 1-11 (Pharmacology)

Section cross-reference(s): 63

FAN.CNT 3

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	US 2002037843	A1	20020328	US 1998-177413	19981023 <--
	US 6514686	B2	20030204		
	WO 2000024390	A1	20000504	WO 1999-US23885	19991014 <--
	W:				
	AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
	RW:				
	GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
	EP 1123090	A1	20010816	EP 1999-954894	19991014 <--
	R:				
	AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO				
	JP 2002528411	T2	20020903	JP 2000-578000	19991014 <--
PRAI	US 1997-847616	B2	19970428		<--
	US 1998-67523	B2	19980428		<--
	US 1998-177413	A2	19981023		<--
	WO 1999-US23885	W	19991014		<--

AB Methods for modulating amyloid deposition in a subject are

described. An effective amt. of at least one ATP binding cassette (ABC) transporter blocker is administered to a subject, such that modulation of **amyloid** deposition occurs. Methods also include administering and effective amt. of at least one ABC transporter blocker, or a pharmaceutically acceptable salt thereof, to a subject such that a disease state assocd. with **amyloidosis** is treated. Packaged pharmaceutical compns. for treating **amyloidosis** are described. The package includes a container for holding an effective amt. of a pharmaceutical compn. and instructions for using the pharmaceutical compn. for treatment of **amyloidosis**. The pharmaceutical compn. includes at least one ABC blocker for modulating **amyloid** deposition in a subject. Methods for identifying agents which modulate **amyloid** deposition in a subject are also described. An effective amt. of at least one ATP binding cassette (ABC) transporter blocker is administered to an organism, such that modulation of **amyloid** deposition occurs.

- ST modulating **amyloidosis**; **amyloid** deposition modulation  
ATP binding cassette transporter blocker
- IT **Brain**  
(ABC transporter blockade in; ATP binding cassette (ABC) transporter blocker for modulating **amyloidosis**)
- IT EST (expressed sequence tag)  
RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(ABC transporter blocker-encoding; ATP binding cassette (ABC) transporter blocker for modulating **amyloidosis**)
- IT Gene, animal  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(ABC1, blockers; ATP binding cassette (ABC) transporter blocker for modulating **amyloidosis**)
- IT Gene, animal  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(ABC2, blockers; ATP binding cassette (ABC) transporter blocker for modulating **amyloidosis**)
- IT Gene, animal  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(ABC3, blockers; ATP binding cassette (ABC) transporter blocker for modulating **amyloidosis**)
- IT Gene, animal  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(ABC7, blockers; ATP binding cassette (ABC) transporter blocker for modulating **amyloidosis**)
- IT Gene, animal  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(ABC8, blockers; ATP binding cassette (ABC) transporter blocker for modulating **amyloidosis**)
- IT **Alzheimer's disease**  
**Amyloidosis**  
**Anti-Alzheimer's agents**  
Bilayer membranes  
Cell membrane  
Drug screening  
Human  
Liposomes  
Membrane, biological  
Multidrug resistance  
(ATP binding cassette (ABC) transporter blocker for modulating **amyloidosis**)
- IT Transport proteins  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(ATP binding cassette (ABC) transporter, blockers; ATP binding cassette (ABC) transporter blocker for modulating **amyloidosis**)
- IT Gene, animal

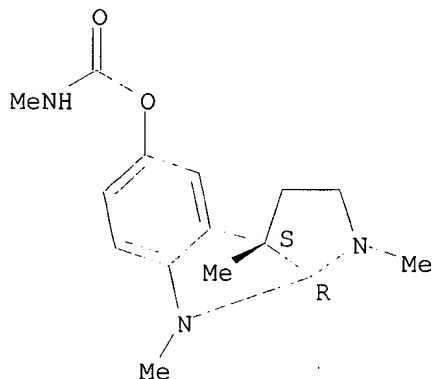
- RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(MDR1, blockers; ATP binding cassette (ABC) transporter blocker for  
modulating **amyloidosis**)
- IT Gene, animal  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(MRP4, inhibitors; ATP binding cassette (ABC) transporter blocker for  
modulating **amyloidosis**)
- IT Gene, animal  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(MRP5, inhibitors; ATP binding cassette (ABC) transporter blocker for  
modulating **amyloidosis**)
- IT P-glycoproteins  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(blockers; ATP binding cassette (ABC) transporter blocker for  
modulating **amyloidosis**)
- IT **Amyloid precursor proteins**  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(cleavage of; ATP binding cassette (ABC) transporter blocker for  
modulating **amyloidosis**)
- IT **Amyloid**  
RL: ADV (Adverse effect, including toxicity); BSU (Biological study,  
unclassified); BIOL (Biological study)  
(deposition of; ATP binding cassette (ABC) transporter blocker for  
modulating **amyloidosis**)
- IT Biological transport  
(efflux, pump; ATP binding cassette (ABC) transporter blocker for  
modulating **amyloidosis**)
- IT Labels  
(for drug packaging; ATP binding cassette (ABC) transporter blocker for  
modulating **amyloidosis**)
- IT Packaging materials  
(for drugs; ATP binding cassette (ABC) transporter blocker for  
modulating **amyloidosis**)
- IT Head  
(injury; ATP binding cassette (ABC) transporter blocker for modulating  
**amyloidosis**)
- IT Gene, animal  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(mdr3, blockers; ATP binding cassette (ABC) transporter blocker for  
modulating **amyloidosis**)
- IT Blood vessel  
(microvessel, of brain, ABC transporter blockade in; ATP binding  
cassette (ABC) transporter blocker for modulating **amyloidosis**  
)
- IT Protein degradation  
(of **amyloid** precursor protein; ATP binding cassette (ABC)  
transporter blocker for modulating **amyloidosis**)
- IT Transport proteins  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(phospholipid-transporting, blockers; ATP binding cassette (ABC)  
transporter blocker for modulating **amyloidosis**)
- IT **Brain, disease**  
(**stroke**; ATP binding cassette (ABC) transporter blocker for  
modulating **amyloidosis**)
- IT **Amyloid**  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(.beta.-; ATP binding cassette (ABC) transporter blocker for modulating  
**amyloidosis**)
- IT 50-53-3, ~~Chlorpromazine~~, biological studies 50-55-5, Reserpine  
51-55-8, ~~Atropine~~, biological studies 52-53-9, Verapamil 54-05-7,  
~~Chloroquine~~ 57-47-6, ~~Physostigmine~~ 61-54-1, Tryptamine  
65-61-2, ~~Acridine orange~~ 83-89-6, Quinacrine 90-34-6, Primaquine  
117-89-5, Trifluoperazine 130-95-0, Quinine 146-48-5, Yohimbine



260-94-6, Acridine 483-10-3, Corynanthine 485-71-2, Cinchonidine  
 525-66-6, Propranolol 10540-29-1, Tamoxifen 59865-13-3, Cyclosporin  
 59865-15-5, Cyclosporin A, 6-[(2S,3R,4R)-3-hydroxy-4-methyl-2-(  
 (methylamino)octanoic acid]- 64657-18-7, 1,9-Dideoxyforskolin  
 66575-29-9, Forskolin 84371-65-3, RU-486 92302-55-1,  
 Benzeneacetonitrile, 3,4-dimethoxy-.alpha.-[3-[[2-(3-  
 methoxyphenyl)ethyl]methylamino]propyl]-.alpha.-(1-methylethyl)- 104987-  
 11-3, FK-506 119666-09-0, AHC-52 121584-18-7, PSC-833 129716-45-6,  
 MS-073 140945-01-3, S9788 143664-11-3, GF120918 158681-49-3, MS-209  
 159997-94-1, VX-710 190454-58-1, VX-853  
 RL: BSU (Biological study, unclassified); PAC (Pharmacological activity);  
**THU (Therapeutic use)**; BIOL (Biological study); USES (Uses)  
 (ATP binding cassette (ABC) transporter blocker for modulating  
**amyloidosis**)

- IT 123955-65-7, RU 49953  
 RL: BSU (Biological study, unclassified); PAC (Pharmacological activity);  
**THU (Therapeutic use)**; BIOL (Biological study); USES (Uses)  
 (RU 49953; ATP binding cassette (ABC) transporter blocker for  
 modulating **amyloidosis**)
- IT 180422-22-4, XR 9051  
 RL: BSU (Biological study, unclassified); PAC (Pharmacological activity);  
**THU (Therapeutic use)**; BIOL (Biological study); USES (Uses)  
 (XR 9051; ATP binding cassette (ABC) transporter blocker for modulating  
**amyloidosis**)
- IT 56-65-5, 5'-ATP, biological studies  
 RL: BSU (Biological study, unclassified); BIOL (Biological study)  
 (cassette binding; ATP binding cassette (ABC) transporter blocker for  
 modulating **amyloidosis**)
- IT **57-47-6**, Physostigmine  
 RL: BSU (Biological study, unclassified); PAC (Pharmacological activity);  
**THU (Therapeutic use)**; BIOL (Biological study); USES (Uses)  
 (ATP binding cassette (ABC) transporter blocker for modulating  
**amyloidosis**)
- RN 57-47-6 HCAPLUS  
 CN Pyrrolo[2,3-b]indol-5-ol, 1,2,3,3a,8,8a-hexahydro-1,3a,8-trimethyl-,  
 methylcarbamate (ester), (3aS,8aR)- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).



L127 ANSWER 8 OF 28 HCAPLUS COPYRIGHT 2003 ACS  
 AN **2002:10812** HCAPLUS  
 DN **136:79718**  
 TI Rapid and sensitive detection of aberrant protein(fibril)  
**aggregation in neurodegenerative disease diagnosis and**  
**drug screening**  
 IN **Bamdad, Cynthia C.; Bamdad, R. Shoshana**

PA **Minerva Biotechnologies Corporation, USA**

SO PCT Int. Appl., 139 pp.

CODEN: PIXXD2

DT Patent

LA English

IC ICM G01N033-68

CC 1-1 (Pharmacology)

Section cross-reference(s): 14

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2002001230	A2	20020103	WO 2001-US20232	20010625
	W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG AU 2001070157 A5 20020108 AU 2001-70157 20010625				
PRAI	US 2000-602689	A	20000623		
	US 2000-631818	A	20000803		
	WO 2001-US20232	W	20010625		
AB.	Methods, assays, and components are described in which biol. samples can be rapidly and sensitively analyzed for the presence of species assocd. with <b>neurodegenerative</b> disease. Techniques and components are provided for diagnosis of disease, as well as for screening of candidate drugs for treatment of <b>neurodegenerative</b> disease. The techniques are simple, extremely sensitive, and utilize readily-available components. Binding species, capable of binding a <b>neurodegenerative</b> disease <b>aggregate</b> -forming or <b>aggregate</b> -forming species, are fastened to surfaces of electrodes and surfaces of particles, or provided free in soln., to bind <b>aggregate</b> -forming species and/or be involved in <b>aggregation</b> .				
ST	aberrant protein fibril <b>aggregation</b> colloid; drug screening <b>neurodegenerative</b> disease kit				
IT	<b>Brain, disease</b> <b>Prion diseases</b> (Creutzfeldt-Jakob; rapid and sensitive detection of aberrant protein(fibril) <b>aggregation</b> in <b>neurodegenerative</b> disease diagnosis and drug screening)				
IT	<b>Prion proteins</b> RL: DGN (Diagnostic use); PRP (Properties); BIOL (Biological study); USES (Uses) (PrPSc; rapid and sensitive detection of aberrant protein(fibril) <b>aggregation</b> in <b>neurodegenerative</b> disease diagnosis and drug screening)				
IT	<b>Voltammetry</b> (a.c.; rapid and sensitive detection of aberrant protein(fibril) <b>aggregation</b> in <b>neurodegenerative</b> disease diagnosis and drug screening)				
IT	<b>Spheres</b> (beads; rapid and sensitive detection of aberrant protein(fibril) <b>aggregation</b> in <b>neurodegenerative</b> disease diagnosis and drug screening)				
IT	<b>Prion proteins</b> RL: DGN (Diagnostic use); PRP (Properties); BIOL (Biological study); USES (Uses) (bovine spongiform encephalopathy; rapid and sensitive detection of aberrant protein(fibril) <b>aggregation</b> in				

- neurodegenerative disease diagnosis and drug screening)**
- IT Proteins  
 RL: DGN (Diagnostic use); PRP (Properties); BIOL (Biological study); USES (Uses)  
 (complexes; rapid and sensitive detection of aberrant protein(fibril) **aggregation in neurodegenerative disease diagnosis and drug screening)**
- IT **Nervous system**  
 (degeneration; rapid and sensitive detection of aberrant protein(fibril) **aggregation in neurodegenerative disease diagnosis and drug screening)**
- IT Self-assembled monolayers  
 (electroactive; rapid and sensitive detection of aberrant protein(fibril) **aggregation in neurodegenerative disease diagnosis and drug screening)**
- IT Immunoassay  
 (enzyme-linked immunosorbent assay; rapid and sensitive detection of aberrant protein(fibril) **aggregation in neurodegenerative disease diagnosis and drug screening)**
- IT Enzymes, biological studies  
 RL: PAC (Pharmacological activity); PRP (Properties); BIOL (Biological study)  
 (inhibitors, capsase; rapid and sensitive detection of aberrant protein(fibril) **aggregation in neurodegenerative disease diagnosis and drug screening)**
- IT Carboxyl group  
 (ionized; rapid and sensitive detection of aberrant protein(fibril) **aggregation in neurodegenerative disease diagnosis and drug screening)**
- IT **Aggregation**  
**Alzheimer's disease**  
 Animal  
 Animal cell  
 Blood analysis  
 Cerebrospinal fluid  
 Colloids  
 Diagnosis  
 Drug screening  
 Feed  
 Fibril  
 High throughput screening  
 Human  
 Immobilization, molecular  
 Livestock  
 Magnetic particles  
 Microtiter plates  
 Milk  
 Molecular association  
 Molecular recognition  
**Parkinson's disease**  
 Protein sequences  
 Test kits  
 Transplant and Transplantation  
 UV and visible spectroscopy  
 (rapid and sensitive detection of aberrant protein(fibril) **aggregation in neurodegenerative disease diagnosis and drug screening)**
- IT Enzymes, biological studies  
 RL: BSU (Biological study, unclassified); PAC (Pharmacological activity); PRP (Properties); BIOL (Biological study)  
 (rapid and sensitive detection of aberrant protein(fibril) **aggregation in neurodegenerative disease diagnosis and drug screening)**

- IT p53 (protein)  
 RL: BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study)  
 (rapid and sensitive detection of aberrant protein(fibril) **aggregation** in **neurodegenerative** disease diagnosis and drug screening)
- IT Metalloenes  
 RL: CPS (Chemical process); PEP (Physical, engineering or chemical process); PRP (Properties); PROC (Process)  
 (rapid and sensitive detection of aberrant protein(fibril) **aggregation** in **neurodegenerative** disease diagnosis and drug screening)
- IT DNA  
 RL: DGN (Diagnostic use); PRP (Properties); BIOL (Biological study); USES (Uses)  
 (rapid and sensitive detection of aberrant protein(fibril) **aggregation** in **neurodegenerative** disease diagnosis and drug screening)
- IT Peptides, biological studies  
 RL: PAC (Pharmacological activity); PRP (Properties); BIOL (Biological study)  
 (rapid and sensitive detection of aberrant protein(fibril) **aggregation** in **neurodegenerative** disease diagnosis and drug screening)
- IT Nucleic acids  
 Oligonucleotides  
 Proteins  
 RL: PRP (Properties)  
 (rapid and sensitive detection of aberrant protein(fibril) **aggregation** in **neurodegenerative** disease diagnosis and drug screening)
- IT Antibodies  
 RL: PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (rapid and sensitive detection of aberrant protein(fibril) **aggregation** in **neurodegenerative** disease diagnosis and drug screening)
- IT **Brain, disease**  
 (**spongiform encephalopathy**, transmissible; rapid and sensitive detection of aberrant protein(fibril) **aggregation** in **neurodegenerative** disease diagnosis and drug screening)
- IT Sensors  
 (surface plasmon resonance chip; rapid and sensitive detection of aberrant protein(fibril) **aggregation** in **neurodegenerative** disease diagnosis and drug screening)
- IT Transferrins  
 RL: BSU (Biological study, unclassified); CPS (Chemical process); PEP (Physical, engineering or chemical process); PRP (Properties); BIOL (Biological study); PROC (Process)  
 (.tau.-transferrins; rapid and sensitive detection of aberrant protein(fibril) **aggregation** in **neurodegenerative** disease diagnosis and drug screening)
- IT **Amyloid**  
 RL: BSU (Biological study, unclassified); PAC (Pharmacological activity); PRP (Properties); BIOL (Biological study)  
 (.beta.-, C-terminal fragment; rapid and sensitive detection of aberrant protein(fibril) **aggregation** in **neurodegenerative** disease diagnosis and drug screening)
- IT **Amyloid**  
 RL: BSU (Biological study, unclassified); PAC (Pharmacological activity); PRP (Properties); BIOL (Biological study)  
 (.beta.-; rapid and sensitive detection of aberrant protein(fibril) **aggregation** in **neurodegenerative** disease diagnosis and drug screening)

and drug screening)  
 IT 167396-02-3 286411-43-6 286411-44-7 286411-46-9 286411-47-0  
 286411-48-1  
 RL: PRP (Properties)  
 (Unclaimed; rapid and sensitive detection of aberrant protein(fibril)  
**aggregation in neurodegenerative** disease diagnosis  
 and drug screening)  
 IT 7732-18-5, Water, biological studies  
 RL: BSU (Biological study, unclassified); PRP (Properties); BIOL  
 (Biological study)  
 (rapid and sensitive detection of aberrant protein(fibril)  
**aggregation in neurodegenerative** disease diagnosis  
 and drug screening)  
 IT 58-85-5, 1H-Thieno[3,4-d]imidazole-4-pentanoic acid, hexahydro-2-oxo-,  
 (3aS,4S,6aR)- 70-18-8, Glycine, L-.gamma.-glutamyl-L-cysteinyl-,  
 properties 102-54-5, Ferrocene 139-13-9, Glycine, N,N-  
 bis(carboxymethyl)- 573-58-0, 1-Naphthalenesulfonic acid,  
 3,3'-[[1,1'-biphenyl]-4,4'-diylbis(azo)]bis4-amino-, disodium salt  
 2390-54-7, Benzothiazolium, 2-[4-(dimethylamino)phenyl]-3,6-dimethyl-,  
 chloride 6066-82-6, 2,5-Pyrrolidinedione, 1-hydroxy- 9001-78-9,  
 Phosphatase, alkaline 9013-20-1, Streptavidin 10487-90-8, Phenol,  
 2,2'-[(6,6'-dimethyl[1,1'-biphenyl]-2,2'-diyl)bis(nitrilomethylidyne)]bis-  
 64691-70-9, Pyridine, 2,2'-[1,2-ethanediylbis(thio-2,1-ethanediyl)]bis-  
 RL: CPS (Chemical process); PEP (Physical, engineering or chemical  
 process); PRP (Properties); PROC (Process)  
 (rapid and sensitive detection of aberrant protein(fibril)  
**aggregation in neurodegenerative** disease diagnosis  
 and drug screening)  
 IT 7440-57-5, Gold, properties  
 RL: DEV (Device component use); PRP (Properties); USES (Uses)  
 (rapid and sensitive detection of aberrant protein(fibril)  
**aggregation in neurodegenerative** disease diagnosis  
 and drug screening)  
 IT 78990-62-2, Calpain 158736-49-3, .beta.-Secretase 338454-52-7,  
 .gamma.-Secretase  
 RL: PAC (Pharmacological activity); PRP (Properties); BIOL (Biological  
 study)  
 (rapid and sensitive detection of aberrant protein(fibril)  
**aggregation in neurodegenerative** disease diagnosis  
 and drug screening)

L127 ANSWER 9 OF 28 HCAPLUS COPYRIGHT 2003 ACS

AN 2001:798758 HCAPLUS

DN 135:339282

TI Nicotine receptor partial agonist, cholinesterase inhibitor, and  
 estrogenic agent composition for treatment of diseases of  
**cognitive** dysfunction in a mammal

IN Coe, Jotham Wadsworth; Sands, Steven Bradley; Harrigan, Edmund Patrick;  
 O'Neill, Brian Thomas; Watsky, Eric Jacob

PA USA

SO U.S. Pat. Appl. Publ., 20 pp.  
 CODEN: USXXCO

DT **Patent**

LA English

IC ICM A61K031-44

ICS A01N043-42

NCL 514299000

CC 1-11 (Pharmacology)

Section cross-reference(s): 63

FAN.CNT 1

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 2001036949	A1	20011101	US 2001-760966	20010116 <--

WO 2001085145 A2 20011115 WO 2001-IB681 20010424 <--  
 WO 2001085145 A3 20020613  
 W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,  
 CO, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM,  
 HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS,  
 LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO,  
 RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ,  
 VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM  
 RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,  
 DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,  
 BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG  
 EP 1280554 A2 20030205 EP 2001-921733 20010424 <--  
 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,  
 IE, SI, LT, LV, FI, RO, MK, CY, AL, TR

PRAI US 2000-202799P P 20000509 <--  
 WO 2001-IB681 W 20010424

AB A pharmaceutical compn. and method of treatment of diseases of  
**cognitive** dysfunction in a mammal comprising administration of a  
 nicotine receptor partial agonist or a pharmaceutically acceptable salt  
 thereof; and an acetylcholinesterase inhibitor, butylcholinesterase  
 inhibitor, an estrogenic agent, selective estrogen receptor modulator or  
 muscarinic agonist or a pharmaceutically acceptable salt thereof; and a  
 pharmaceutically acceptable carrier. The nicotine receptor partial  
 agonist and acetylcholinesterase inhibitor, butylcholinesterase inhibitor,  
 estrogen, selective estrogen receptor modulator or muscarinic agonist are  
 present in amts. that render the compn. effective enhancing  
**cognition** or in the treatment of diseases of **cognitive**  
 dysfunction including but not limited to **Alzheimer's Disease**,  
 mild **cognitive** impairment, age-related **cognitive**  
 decline, vascular **dementia**, **Parkinson's disease**  
**dementia**, **Huntington's Disease**, **Stroke**, **TBI**, **AIDS**  
 assocd. **dementia** and schizophrenia. The method of using these  
 compns. is also disclosed.

ST **cognitive** dysfunction treatment compn; nicotinic agonist  
**cognitive** dysfunction treatment compn; cholinesterase inhibitor  
**cognitive** dysfunction treatment compn; estrogen **cognitive**  
 dysfunction treatment compn

IT **Nervous system**  
 (**Huntington's chorea**; nicotine receptor partial  
 agonist, cholinesterase inhibitor, and estrogenic agent compn. for  
 treatment of diseases of **cognitive** dysfunction in a mammal)

IT **Mental disorder**  
 (**cognitive**; nicotine receptor partial agonist, cholinesterase  
 inhibitor, and estrogenic agent compn. for treatment of diseases of  
**cognitive** dysfunction in a mammal)

IT **Mental disorder**  
 (**dementia**; nicotine receptor partial agonist, cholinesterase  
 inhibitor, and estrogenic agent compn. for treatment of diseases of  
**cognitive** dysfunction in a mammal)

IT **Cognition**  
 (disorder; nicotine receptor partial agonist, cholinesterase inhibitor,  
 and estrogenic agent compn. for treatment of diseases of  
**cognitive** dysfunction in a mammal)

IT **Alzheimer's disease**  
 Muscarinic agonists  
 Nicotinic agonists  
**Parkinson's disease**  
**Schizophrenia**  
 (nicotine receptor partial agonist, cholinesterase inhibitor, and  
 estrogenic agent compn. for treatment of diseases of **cognitive**  
 dysfunction in a mammal)

IT Estrogens  
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(nicotine receptor partial agonist, cholinesterase inhibitor, and estrogenic agent compn. for treatment of diseases of **cognitive** dysfunction in a mammal)

IT **Brain, disease**

(**stroke**; nicotine receptor partial agonist, cholinesterase inhibitor, and estrogenic agent compn. for treatment of diseases of **cognitive** dysfunction in a mammal)

IT 9000-81-1, Acetylcholinesterase 9001-08-5

RL: BSU (Biological study, unclassified); BIOL (Biological study)

(inhibitors; nicotine receptor partial agonist, cholinesterase inhibitor, and estrogenic agent compn. for treatment of diseases of **cognitive** dysfunction in a mammal)

IT 50-28-2, Estradiol, biological studies 52-68-6, Metrifonate  
 57-47-6, Physostigmine 59-99-4, Neostigmine 63-75-2, Arecoline  
 70-22-4, Oxotremorine 92-13-7, Pilocarpine 101-26-8, Mestizon  
 321-64-2, Tacrine 357-70-0, Galanthamine 469-22-7, Eseroline  
 10540-29-1, Tamoxifen 69718-72-5 82413-20-5, Droloxifene 84449-90-1,  
 Raloxifene 102518-79-6, Huperzine A 120011-70-3, Aricept  
 123441-03-2, Rivastigmine 131986-45-3, Xanomeline 132236-18-1,  
 Zifrosilone 139314-01-5, Quilostigmine 139886-32-1, Milameline  
 145209-30-9, Tolserine 145209-39-8, Cymserine 145209-50-3,  
 Thiatolserine 145209-51-4, Thiacymseline 145508-78-7, Icopezil  
 159912-53-5, Sabcomeline 180916-16-9, Lasofoxifene 207391-08-0  
 207391-10-4 207391-12-6 207391-15-9 207391-18-2 207391-21-7  
 207391-24-0 207391-27-3 207391-28-4 207391-29-5 207391-34-2  
 207391-36-4 207391-37-5 207391-38-6 207391-40-0 207391-41-1  
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 207391-67-1 207391-74-0 230615-75-5 248275-68-5 248275-79-8  
 248275-81-2 248275-95-8 248276-19-9 249296-44-4 287973-23-3  
 287973-26-6 287973-27-7 328055-76-1 328055-77-2 328055-78-3  
 328055-79-4 328055-80-7 328055-81-8 328055-83-0 328055-84-1  
 328055-85-2 328055-86-3 328055-87-4 328055-88-5 328055-89-6  
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 328056-26-4 328056-27-5 328056-28-6 328056-29-7 328056-30-0  
 328056-66-2 357424-19-2 357424-20-5 371238-38-9 371238-39-0  
 371238-40-3 371238-41-4

RL: **THU (Therapeutic use)**; BIOL (Biological study); USES (Uses)

(nicotine receptor partial agonist, cholinesterase inhibitor, and estrogenic agent compn. for treatment of diseases of **cognitive** dysfunction in a mammal)

IT 57-47-6, Physostigmine

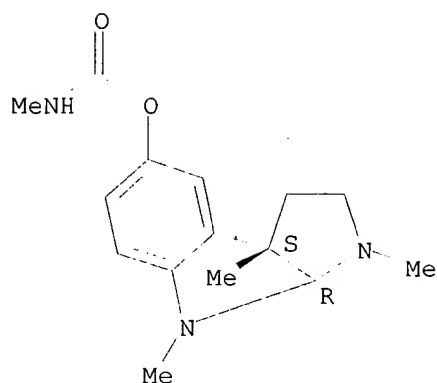
RL: **THU (Therapeutic use)**; BIOL (Biological study); USES (Uses)

(nicotine receptor partial agonist, cholinesterase inhibitor, and estrogenic agent compn. for treatment of diseases of **cognitive** dysfunction in a mammal)

RN 57-47-6 HCAPLUS

CN Pyrrolo[2,3-b]indol-5-ol, 1,2,3,3a,8,8a-hexahydro-1,3a,8-trimethyl-, methylcarbamate (ester), (3aS,8aR)- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).



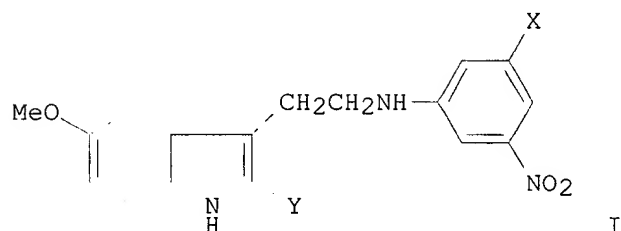
L127 ANSWER 10 OF 28 HCAPLUS COPYRIGHT 2003 ACS  
 AN 2001:792336 HCAPLUS  
 DN 135:339274  
 TI Method for the treatment of **neurological** or  
**neuropsychiatric** disorders with melatonin antagonists  
 IN Willis, Gregory Lynn  
 PA Clarence Pty Ltd., Australia  
 SO U.S., 31 pp., Cont.-in-part of Appl. No. PCT/AU97/00661  
 CODEN: USXXAM  
 DT **Patent**  
 LA English  
 IC ICM A61K031-405  
 NCL 514415000  
 CC 1-11 (Pharmacology)  
 FAN.CNT 4

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	US 6310085	B1	20011030	US 1999-285859	19990402 <--
	WO 9815267	A1	19980416	WO 1997-AU661	19971003 <--
	W:				
	AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE,				
	DK, EE, ES, FI, GB, GE, GH, HU, ID, IL, IS, JP, KE, KG, KP, KR,				
	KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ,				
	PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG,				
	US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
	RW:				
	GH, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, DE, DK, ES, FI, FR,				
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	WO 2000059504	A1	20001012	WO 2000-AU275	20000331 <--
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	SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW,				
	AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
	RW:				
	GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE,				
	DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF,				
	CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
	BR 2000009524	A	20020219	BR 2000-9524	20000331 <--
	EP 1189613	A1	20020327	EP 2000-912271	20000331 <--
	R:				
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	IE, SI, LT, LV, FI, RO				
	JP 2002541105	T2	20021203	JP 2000-609068	20000331 <--
	EE 200100511	A	20021216	EE 2001-511	20000331 <--
	NO 2001004674	A	20010926	NO 2001-4674	20010926 <--
	US 2002068692	A1	20020606	US 2001-971783	20011009 <--



PRAI WO 1997-AU661 A2 19971003 <--  
 AU 1996-2745 A 19961004 <--  
 US 1999-285859 A 19990402 <--  
 WO 2000-AU275 W 20000331 <--

GI



- AB A method for the treatment and/or prophylaxis of a **neurol.** or **neuropsychiatric** disorder assocd. with altered dopamine function comprises administering melatonin antagonists I (X = NO<sub>2</sub>, N<sub>3</sub>; Y = H, I) to a patient in need thereof. I (X = NO<sub>2</sub>; Y = H) (ML-23) prevented the development of severe motor impairment and severe body wt. loss typically exhibited by **neurotoxin** 6-hydroxydopamine-treated rats.
- ST **neurol** disorder **neuropsychiatric** disorder treatment  
melatonin antagonist
- IT **Brain, disease**  
(Gilles de la Tourette syndrome; method for treatment of **neurol.** or **neuropsychiatric** disorders with melatonin antagonists in combination with light therapy)
- IT **Nervous system**  
(Huntington's chorea; method for treatment of **neurol.** or **neuropsychiatric** disorders with melatonin antagonists)
- IT **Mental disorder**  
(Korsakow's syndrome; method for treatment of **neurol.** or **neuropsychiatric** disorders with melatonin antagonists in combination with light therapy)
- IT **Mental disorder**  
(Pick's disease; method for treatment of **neurol.** or **neuropsychiatric** disorders with melatonin antagonists)
- IT **Mental disorder**  
(Punch drunk syndrome; method for treatment of **neurol.** or **neuropsychiatric** disorders with melatonin antagonists)
- IT **Mental disorder**  
(Sundowner's syndrome; method for treatment of **neurol.** or **neuropsychiatric** disorders with melatonin antagonists in combination with light therapy)
- IT **Stress, animal**  
(acute; method for treatment of **neurol.** or **neuropsychiatric** disorders with melatonin antagonists in combination with light therapy)
- IT **Mental disorder**  
(agoraphobia; method for treatment of **neurol.** or **neuropsychiatric** disorders with melatonin antagonists)
- IT **Appetite**  
(anorexia **nervosa**; method for treatment of **neurol.** or **neuropsychiatric** disorders with melatonin antagonists)
- IT **Cachexia**  
(anorexia; method for treatment of **neurol.** or **neuropsychiatric** disorders with melatonin antagonists)
- IT **Anorexia**

- (cachexia; method for treatment of **neurol.** or **neuropsychiatric** disorders with melatonin antagonists in combination with light therapy)
- IT Ion channel blockers  
(calcium; method for treatment of **neurol.** or **neuropsychiatric** disorders with melatonin antagonists in combination with a calcium channel blocker)
- IT Mental disorder  
(dementia; method for treatment of **neurol.** or **neuropsychiatric** disorders with melatonin antagonists in combination with light therapy)
- IT Mental disorder  
(depression, anxiety disorders due to; method for treatment of **neurol.** or **neuropsychiatric** disorders with melatonin antagonists)
- IT Nervous system  
(disease, malignant syndrome; method for treatment of **neurol.** or **neuropsychiatric** disorders with melatonin antagonists)
- IT Nervous system  
(dystonia, acute; method for treatment of **neurol.** or **neuropsychiatric** disorders with melatonin antagonists)
- IT Brain, disease  
(ischemia, transient; method for treatment of **neurol.** or **neuropsychiatric** disorders with melatonin antagonists in combination with light therapy)
- IT Anti-Alzheimer's agents  
Antiparkinsonian agents  
Anxiolytics  
Movement disorders  
Multiple sclerosis  
(method for treatment of **neurol.** or **neuropsychiatric** disorders with melatonin antagonists)
- IT Nervous system agents  
Phototherapy  
Psychotropics  
Schizophrenia  
Wernicke-Korsakoff syndrome  
(method for treatment of **neurol.** or **neuropsychiatric** disorders with melatonin antagonists in combination with light therapy)
- IT Pineal gland  
(method for treatment of **neurol.** or **neuropsychiatric** disorders with melatonin antagonists in combination with surgical ablation or destruction of the pineal gland)
- IT Nervous system  
(multiple system atrophy; method for treatment of **neurol.** or **neuropsychiatric** disorders with melatonin antagonists)
- IT Mental disorder  
(obsession-compulsion; method for treatment of **neurol.** or **neuropsychiatric** disorders with melatonin antagonists)
- IT Anxiety  
(panic disorder; method for treatment of **neurol.** or **neuropsychiatric** disorders with melatonin antagonists)
- IT Movement disorders  
(periodic limb movement disorders; method for treatment of **neurol.** or **neuropsychiatric** disorders with melatonin antagonists in combination with light therapy)
- IT Mental disorder  
(post-traumatic stress disorder; method for treatment of **neurol.** or **neuropsychiatric** disorders with melatonin antagonists)
- IT Mental disorder  
(post-traumatic stress disorder; method for treatment of **neurol.** or **neuropsychiatric** disorders with melatonin antagonists in combination with light therapy)

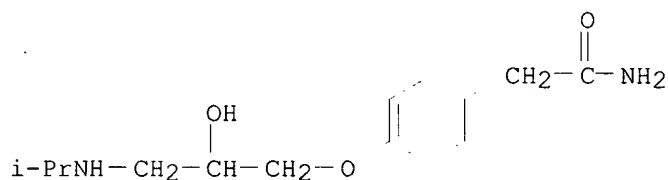
- IT Paralysis  
(progressive subnuclear; method for treatment of **neurol.** or **neuropsychiatric** disorders with melatonin antagonists in combination with light therapy)
- IT Movement disorders  
(restless leg syndrome; method for treatment of **neurol.** or **neuropsychiatric** disorders with melatonin antagonists in combination with light therapy)
- IT Brain, disease  
(**stroke**; method for treatment of **neurol.** or **neuropsychiatric** disorders with melatonin antagonists)
- IT Nervous system  
(tardive dyskinesia; method for treatment of **neurol.** or **neuropsychiatric** disorders with melatonin antagonists)
- IT Adrenoceptor antagonists  
(.beta.-; method for treatment of **neurol.** or **neuropsychiatric** disorders with melatonin antagonists in combination with a .beta.-adrenergic antagonist)
- IT 73-31-4, Melatonin  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(antagonists; **neurol.** or **neuropsychiatric** disorders assocd. with altered dopamine function treatment with melatonin antagonists)
- IT 152302-33-5, S-20928  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(method for treatment of **neurol.** or **neuropsychiatric** disorders with melatonin antagonists)
- IT 115007-18-6, ML-23  
RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(method for treatment of **neurol.** or **neuropsychiatric** disorders with melatonin antagonists)
- IT 9002-79-3, Melanocyte stimulating hormone  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(method for treatment of **neurol.** or **neuropsychiatric** disorders with melatonin antagonists in combination with MSH)
- IT 29122-68-7, Atenolol  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); **THU (Therapeutic use)**; BIOL (Biological study); USES (Uses)  
(method for treatment of **neurol.** or **neuropsychiatric** disorders with melatonin antagonists in combination with a drug that alters dopamine function)
- IT 51-61-6, Dopamine, biological studies  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study)  
(**neurol.** or **neuropsychiatric** disorders assocd. with altered dopamine function treatment with melatonin antagonists)

RE.CNT 20 THERE ARE 20 CITED REFERENCES AVAILABLE FOR THIS RECORD

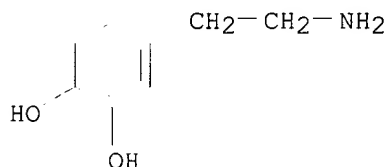
RE

- (1) Alder; Psychopharmacology Bulletin 1991, V27(2), P107
- (2) Anon; EP 0146113 1985 HCAPLUS
- (3) Anon; WO 9529173 1995 HCAPLUS
- (4) Artmenko, A; Arch Neurol 1996, V44
- (5) Chuprikov; US 5137018 1992
- (6) Depreux; US 5552418 1996 HCAPLUS
- (7) Dubocovich; US 5093352 1992 HCAPLUS
- (8) Dubocovich; US 5283343 1994 HCAPLUS
- (9) Horn; US 5071875 1991 HCAPLUS
- (10) Koller; Arch Neurol 1987, V44, P921 MEDLINE

- (11) Martindale; The Extra Pharmacopoeia 28th Edition 1982, P1337  
 (12) Merk Research Laboratories; The Merk Manual of Diagnosis and Therapy, 16th Edition 1992, P1499  
 (13) Miles; Biol Psychiatry 1988, V23, P405 HCAPLUS  
 (14) Sandyk, R; Intern J Neuroscience 1992, V66, P1 MEDLINE  
 (15) Sandyk, R; Intern J Neuroscience 1993, V68, P85 MEDLINE  
 (16) Searfoss; US 5046494 1991  
 (17) Sherer; Neurosci Lett 1985, V58(3), P277 MEDLINE  
 (18) Wilbur; Prog Neuro-Psychopharmacol and Biol Psychiat 1988, V12, P849 MEDLINE  
 (19) Yous; US 5616614 1997 HCAPLUS  
 (20) Zisapel; US 4880826 1989 HCAPLUS
- IT 29122-68-7, Atenolol  
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (method for treatment of **neurol.** or **neuropsychiatric** disorders with melatonin antagonists in combination with a drug that alters dopamine function)
- RN 29122-68-7 HCAPLUS  
 CN Benzeneacetamide, 4-[2-hydroxy-3-[(1-methylethyl)amino]propoxy]- (9CI)  
 (CA INDEX NAME)



- IT 51-61-6, Dopamine, biological studies  
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study)  
 (**neurol.** or **neuropsychiatric** disorders assocd. with altered dopamine function treatment with melatonin antagonists)
- RN 51-61-6 HCAPLUS  
 CN 1,2-Benzenediol, 4-(2-aminoethyl)- (9CI) (CA INDEX NAME)



L127 ANSWER 11 OF 28 HCAPLUS COPYRIGHT 2003 ACS  
 AN 2001:780668 HCAPLUS  
 DN 135:335153  
 TI Treatment of **neurodegenerative** disease  
 IN Bamdad, R. Shoshanna; Bamdad, Cynthia C.  
 PA Minerva Biotechnologies Corporation, USA  
 SO PCT Int. Appl., 82 pp.  
 CODEN: PIXXD2  
 DT Patent  
 LA English  
 IC ICM A61K031-00  
 CC 63-6 (Pharmaceuticals)

Section cross-reference(s): 1

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2001078709	A2	20011025	WO 2001-US12484	20010412 <--
	W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
	US 2003060487	A1	20030327	US 2001-835099	20010412 <--
PRAI	US 2000-196497P	P	20000412 <--		
	US 2000-214221P	P	20000623 <--		
	US 2000-248890P	P	20001115 <--		
OS	MARPAT 135:335153				
AB	<p>The invention relates to treatments for peptide <b>aggregation</b> assocd. with disease states such as <b>neurodegenerative</b> disease, particularly physiol. assocd. with <b>Alzheimer's</b> Disease, and non-<b>neurodegenerative</b> disease <b>aggregation</b>. Other aspects of the invention also provides a variety of novel assays for screening candidate drugs. Yet another aspects of the present invention also provides a series of compns. useful for treatment of <b>neurol.</b> disease as detd. from these assays. These compns. can be packaged in kits. Other aspects of the invention also relate to the use of these compns. for the treatment and/or prevention of patients susceptible to or exhibiting of a disease characteristic of fibril formation or aberrant protein <b>aggregation</b>. Examples are given for monitoring drug activity as a function of time for drug profiling and cell-based screening assay for candidate drugs for affecting <b>aggregate</b> formation at a variety of stages of biochem. progression.</p>				
ST	<b>neurodegenerative</b> disease treatment compn				
IT	<b>Brain, disease</b>				
	<b>Prion diseases</b>				
	(Creutzfeldt-Jakob; treatment of <b>neurodegenerative</b> disease)				
IT	<b>Nervous system</b>				
	(Huntington's chorea; treatment of <b>neurodegenerative</b> disease)				
IT	<b>Nervous system</b>				
	(degeneration; treatment of <b>neurodegenerative</b> disease)				
IT	<b>Mental disorder</b>				
	(dementia; treatment of <b>neurodegenerative</b> disease)				
IT	<b>Amyloidosis</b>				
	(familial <b>amyloidotic polyneuropathy</b> , type IV; treatment of <b>neurodegenerative</b> disease)				
IT	<b>Brain, disease</b>				
	<b>Prion diseases</b>				
	(fatal familial insomnia; treatment of <b>neurodegenerative</b> disease)				
IT	<b>Insomnia</b>				
	(fatal familial; treatment of <b>neurodegenerative</b> disease)				
IT	<b>Brain, disease</b>				
	<b>Prion diseases</b>				
	(kuru; treatment of <b>neurodegenerative</b> disease)				
IT	<b>Nerve, disease</b>				
	(polyneuropathy; treatment of <b>neurodegenerative</b> disease)				
IT	<b>Aggregates</b>				

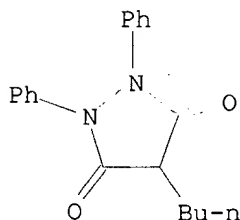
- (protein, formation of; treatment of **neurodegenerative** disease)
- IT **Amyloidosis**  
(senile; treatment of **neurodegenerative** disease)
- IT **Brain, disease**  
(spongiform encephalopathy; treatment of **neurodegenerative** disease)
- IT **Brain, disease**  
(spongiform myeloencephalopathy; treatment of **neurodegenerative** disease)
- IT **Brain, disease**  
(stroke; treatment of **neurodegenerative** disease)
- IT **Alzheimer's disease**  
**Parkinson's disease**  
**Sickle cell anemia**  
(treatment of **neurodegenerative** disease)
- IT **Diabetes mellitus**  
(type II; treatment of **neurodegenerative** disease)
- IT 50-33-9, Phenylbutazone, biological studies 52-01-7, Spironolactone 55-10-7, Vanillylmandelic acid 57-47-6, (-)-Physostigmine 61-76-7, Phenylephrine hydrochloride 65-28-1, Phentolamine mesylate 65-29-2, Gallamine triethiodide 80-77-3, Chlormezanone 102-02-3, 1-Phenylbiguanide 125-33-7, Primidone 130-61-0, Thioridazine hydrochloride 136-47-0, Tetracaine hydrochloride 146-56-5, FLuphenazine dihydrochloride 300-08-3, Arecoline hydrobromide 504-24-5, 4-Aminopyridine 518-28-5, Podophyllotoxin 581-88-4, Debrisoquin sulfate 614-39-1, Procainamide hydrochloride 770-05-8, Octopamine hydrochloride 1011-74-1, Normetanephrine hydrochloride 1069-66-5, Sodium valproate 1867-73-8 1952-15-4 2145-56-4 4789-68-8, Octoclotheptin maleate 13153-27-0, S-(4-Nitrobenzyl)-6-thioguanosine 13523-86-9, Pindolol 15307-79-6, Diclofenac sodium 15676-16-1, Sulpiride 16709-43-6, cis-Dioxolane 17560-51-9, Metolazone 27833-64-3, Loxapine succinate 29122-68-7, Atenolol 30817-59-5, Oxotremorine methiodide 34661-85-3, 5-Methylurapidil 35873-49-5, 8-Cyclopentyl-1,3-dimethylxanthine 38048-32-7 41094-88-6, Tracazolate 51481-61-9, Cimetidine 53296-10-9, 2-Phenylaminoadenosine 56715-13-0, 64710-63-0 75614-89-0 89805-39-0 93379-54-5, S-Atenolol 96850-13-4 100069-68-9 104809-20-3 109028-10-6, CGS-12066B 123064-80-2 127299-93-8 147416-96-4, Telenzepine dihydrochloride 148440-81-7 149981-25-9 153587-01-0 175615-76-6 192575-19-2 369647-58-5 369647-59-6 369647-60-9
- RL: THU (**Therapeutic use**); BIOL (Biological study); USES (Uses)  
(treatment of **neurodegenerative** disease)
- IT 50-33-9, Phenylbutazone, biological studies 52-01-7, Spironolactone 55-10-7, Vanillylmandelic acid 57-47-6, (-)-Physostigmine 61-76-7, Phenylephrine hydrochloride 65-28-1, Phentolamine mesylate 65-29-2, Gallamine triethiodide 80-77-3, Chlormezanone 102-02-3, 1-Phenylbiguanide 125-33-7, Primidone 130-61-0, Thioridazine hydrochloride 136-47-0, Tetracaine hydrochloride 146-56-5, FLuphenazine dihydrochloride 300-08-3, Arecoline hydrobromide 504-24-5, 4-Aminopyridine 518-28-5, Podophyllotoxin 581-88-4, Debrisoquin sulfate 614-39-1, Procainamide hydrochloride 770-05-8, Octopamine hydrochloride 1011-74-1, Normetanephrine hydrochloride 1069-66-5, Sodium valproate 1867-73-8

1952-15-4 2145-56-4 4789-68-8, Octoclotheptin  
 maleate 13153-27-0, S-(4-Nitrobenzyl)-6-thioguanosine  
 13523-86-9, Pindolol 15307-79-6, Diclofenac sodium  
 15676-16-1, Sulpiride 16709-43-6, cis-Dioxolane  
 17560-51-9, Metolazone 27833-64-3, Loxapine succinate  
 29122-68-7, Atenolol 30817-59-5, Oxótremorine methiodide  
 34661-85-3, 5-Methylurapidil 35873-49-5,  
 8-Cyclopentyl-1,3-dimethylxanthine 38048-32-7 41094-88-6  
 , Tracazolate 51481-61-9, Cimetidine 53296-10-9,  
 2-Phenylaminoadenosine 56715-13-0 64710-63-0  
 75614-89-0 89805-39-0 93379-54-5, S-Atenolol  
 96850-13-4 100069-68-9 104809-20-3  
 109028-10-6, CGS-12066B 123064-80-2 127299-93-8  
 147416-96-4, Telenzepine dihydrochloride 148440-81-7  
 149981-25-9 153587-01-0 175615-76-6  
 192575-19-2 369647-58-5 369647-59-6  
 369647-60-9

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (treatment of neurodegenerative disease)

RN 50-33-9 HCAPLUS

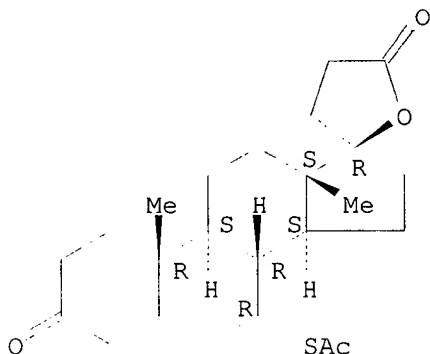
CN 3,5-Pyrazolidinedione, 4-butyl-1,2-diphenyl- (6CI, 8CI, 9CI) (CA INDEX NAME)



RN 52-01-7 HCAPLUS

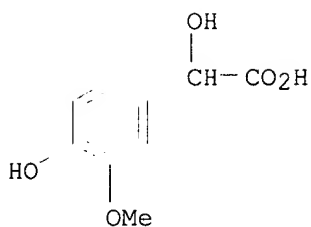
CN Pregn-4-ene-21-carboxylic acid, 7-(acetylthio)-17-hydroxy-3-oxo-,  
 .gamma.-lactone, (7.alpha.,17.alpha.)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



RN 55-10-7 HCAPLUS

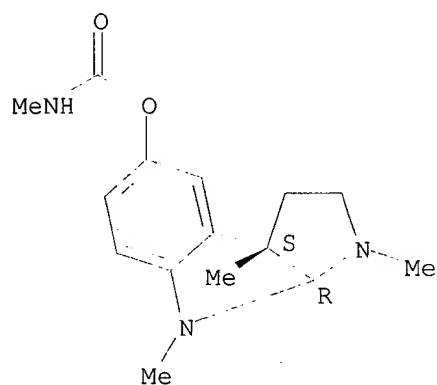
CN Benzeneacetic acid, .alpha.,4-dihydroxy-3-methoxy- (9CI) (CA INDEX NAME)



RN 57-47-6 HCAPLUS

CN Pyrrolo[2,3-b]indol-5-ol, 1,2,3,3a,8,8a-hexahydro-1,3a,8-trimethyl-, methylcarbamate (ester), (3aS,8aR)- (9CI) (CA INDEX NAME)

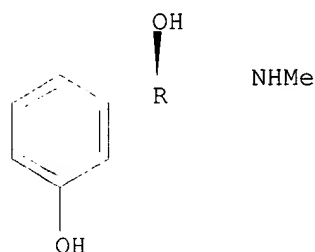
Absolute stereochemistry. Rotation (-).



RN 61-76-7 HCAPLUS

CN Benzenemethanol, 3-hydroxy-.alpha.-[(methylamino)methyl]-, hydrochloride, (.alpha.R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



● HCl

RN 65-28-1 HCAPLUS

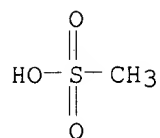
CN Phenol, 3-[[[4,5-dihydro-1H-imidazol-2-yl)methyl](4-methylphenyl)amino]-, monomethanesulfonate (salt) (9CI) (CA INDEX NAME)

CM 1

CRN 75-75-2

CMF C H4 O3 S

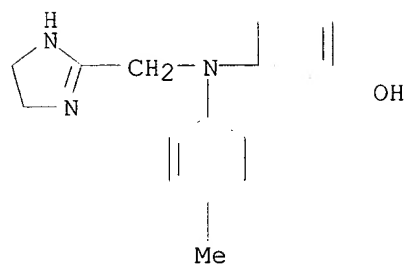




CM 2

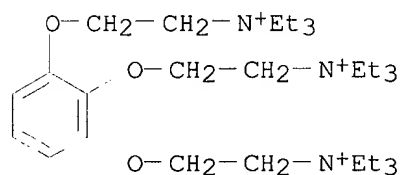
CRN 50-60-2

CMF C17 H19 N3 O



RN 65-29-2 HCAPLUS

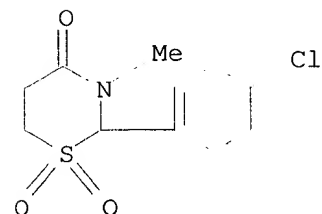
CN Ethanaminium, 2,2',2''-[1,2,3-benzenetriyltris(oxy)]tris[N,N,N-triethyl-, triiodide (9CI) (CA INDEX NAME)



● 3 I<sup>-</sup>

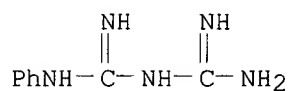
RN 80-77-3 HCAPLUS

CN 4H-1,3-Thiazin-4-one, 2-(4-chlorophenyl)tetrahydro-3-methyl-, 1,1-dioxide (9CI) (CA INDEX NAME)



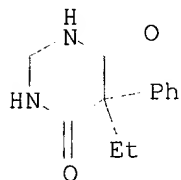
RN 102-02-3 HCAPLUS

CN Imidodicarbonimidic diamide, N-phenyl- (9CI) (CA INDEX NAME)



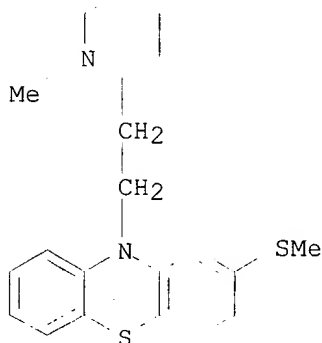
RN 125-33-7 HCAPLUS

CN 4,6(1H,5H)-Pyrimidinedione, 5-ethyldihydro-5-phenyl- (6CI, 8CI, 9CI) (CA INDEX NAME)



RN 130-61-0 HCAPLUS

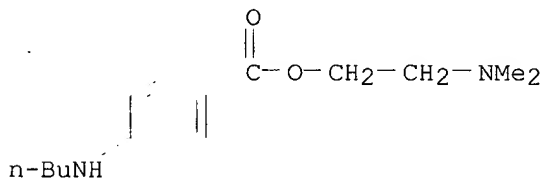
CN 10H-Phenothiazine, 10-[2-(1-methyl-2-piperidinyl)ethyl]-2-(methylthio)-, monohydrochloride (9CI) (CA INDEX NAME)



● HCl

RN 136-47-0 HCAPLUS

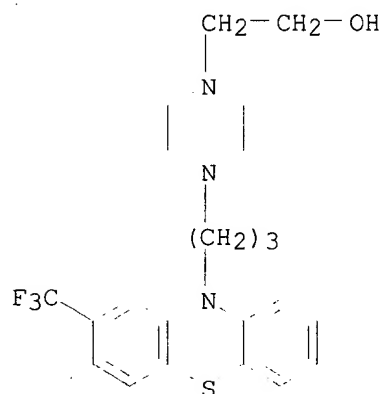
CN Benzoic acid, 4-(butylamino)-, 2-(dimethylamino)ethyl ester, monohydrochloride (9CI) (CA INDEX NAME)



● HCl

RN 146-56-5 HCAPLUS

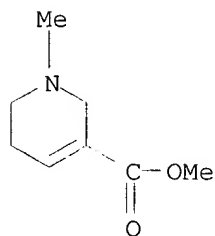
CN 1-Piperazineethanol, 4-[3-[2-(trifluoromethyl)-10H-phenothiazin-10-yl]propyl]-, dihydrochloride (9CI) (CA INDEX NAME)



● 2 HCl

RN 300-08-3 HCAPLUS

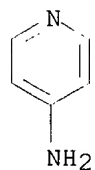
CN 3-Pyridinecarboxylic acid, 1,2,5,6-tetrahydro-1-methyl-, methyl ester, hydrobromide (9CI) (CA INDEX NAME)



● HBr

RN 504-24-5 HCAPLUS

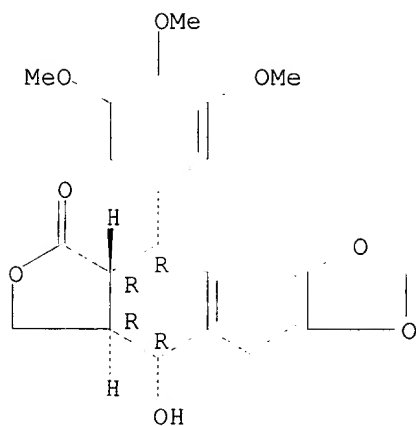
CN 4-Pyridinamine (9CI) (CA INDEX NAME)



RN 518-28-5 HCAPLUS

CN Furo[3',4':6,7]naphtho[2,3-d]-1,3-dioxol-6(5aH)-one, 5,8,8a,9-tetrahydro-9-hydroxy-5-(3,4,5-trimethoxyphenyl)-, (5R,5aR,8aR,9R)- (9CI) (CA INDEX NAME)

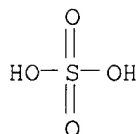
Absolute stereochemistry. Rotation (-).



RN 581-88-4 HCAPLUS  
 CN 2(1H)-Isoquinolinecarboximidamide, 3,4-dihydro-, sulfate (2:1) (9CI) (CA INDEX NAME)

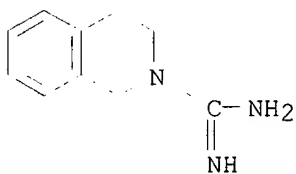
CM 1

CRN 7664-93-9  
 CMF H2 O4 S

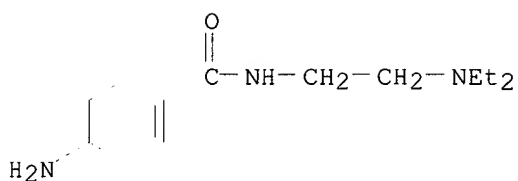


CM 2

CRN 1131-64-2  
 CMF C10 H13 N3

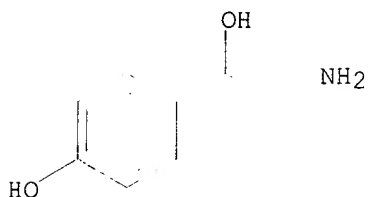


RN 614-39-1 HCAPLUS  
 CN Benzamide, 4-amino-N-[2-(diethylamino)ethyl]-, monohydrochloride (9CI) (CA INDEX NAME)



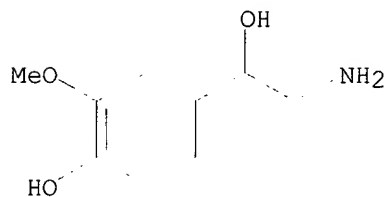
● HCl

RN 770-05-8 HCAPLUS  
 CN Benzenemethanol, .alpha.-(aminomethyl)-4-hydroxy-, hydrochloride (9CI)  
 (CA INDEX NAME)



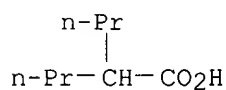
● HCl

RN 1011-74-1 HCAPLUS  
 CN Benzenemethanol, .alpha.-(aminomethyl)-4-hydroxy-3-methoxy-, hydrochloride  
 (9CI) (CA INDEX NAME)



● HCl

RN 1069-66-5 HCAPLUS  
 CN Pentanoic acid, 2-propyl-, sodium salt (9CI) (CA INDEX NAME)

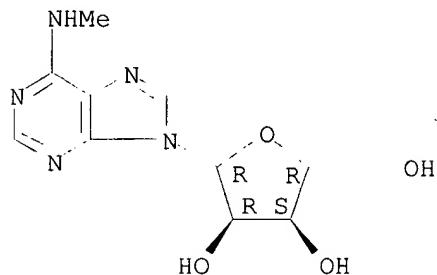


● Na

RN 1867-73-8 HCAPLUS

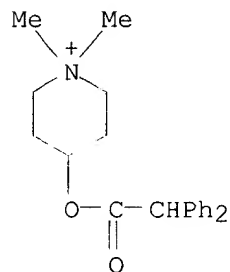
CN Adenosine, N-methyl- (6CI, 7CI, 8CI, 9CI) (CA INDEX NAME)

Absolute stereochemistry.



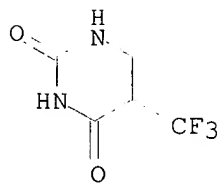
RN 1952-15-4 HCAPLUS

CN Piperidinium, 4-[(diphenylacetyl)oxy]-1,1-dimethyl-, iodide (9CI) (CA INDEX NAME)

● I<sup>-</sup>

RN 2145-56-4 HCAPLUS

CN 2,4(1H,3H)-Pyrimidinedione, dihydro-5-(trifluoromethyl)- (9CI) (CA INDEX NAME)



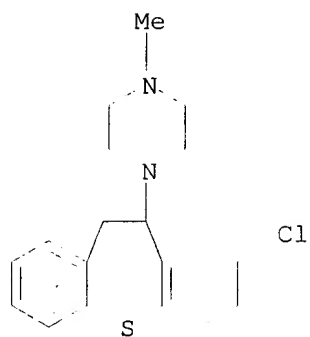
RN 4789-68-8 HCAPLUS

CN Piperazine, 1-(8-chloro-10,11-dihydrodibenzo[b,f]thiepin-10-yl)-4-methyl-, (2Z)-2-butenedioate (1:1) (9CI) (CA INDEX NAME)

CM 1

CRN 13448-22-1

CMF C19 H21 Cl N2 S

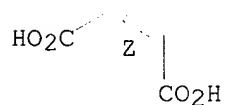


CM 2

CRN 110-16-7

CMF C4 H4 O4

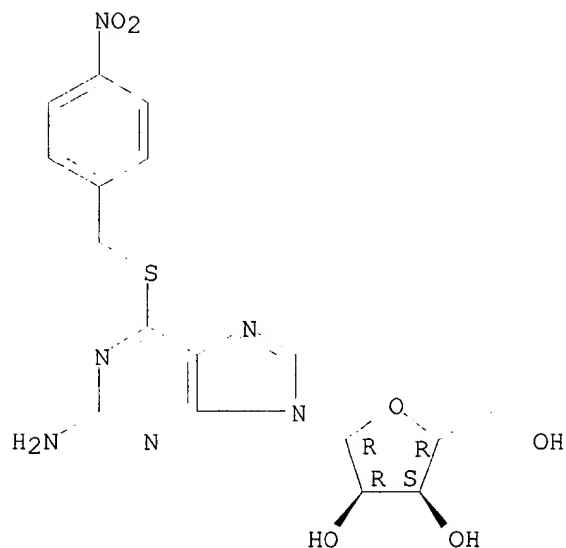
Double bond geometry as shown.



RN 13153-27-0 HCAPLUS

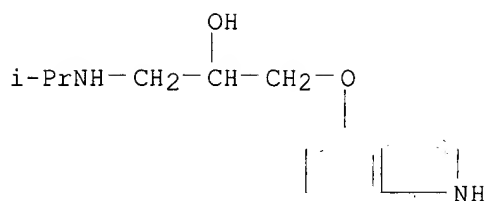
CN Guanosine, 6-S-[(4-nitrophenyl)methyl]-6-thio- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

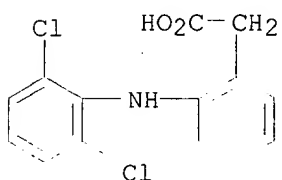


RN 13523-86-9 HCAPLUS

CN 2-Propanol, 1-(1H-indol-4-yloxy)-3-[(1-methylethyl)amino]- (9CI) (CA INDEX NAME)

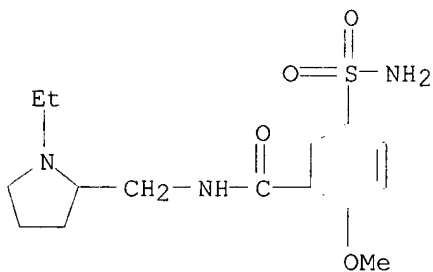


RN 15307-79-6 HCAPLUS  
 CN Benzeneacetic acid, 2-[(2,6-dichlorophenyl)amino]-, monosodium salt (9CI)  
 (CA INDEX NAME)



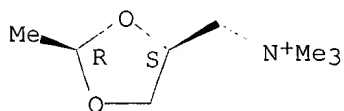
● Na

RN 15676-16-1 HCAPLUS  
 CN Benzamide, 5-(aminosulfonyl)-N-[(1-ethyl-2-pyrrolidinyl)methyl]-2-methoxy-  
 (9CI) (CA INDEX NAME)



RN 16709-43-6 HCAPLUS  
 CN 1,3-Dioxolane-4-methanaminium, N,N,N,2-tetramethyl-, iodide, (2R,4S)-rel-  
 (9CI) (CA INDEX NAME)

Relative stereochemistry.

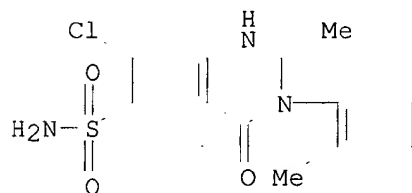


● I<sup>-</sup>



RN 17560-51-9 HCAPLUS

CN 6-Quinazolinesulfonamide; 7-chloro-1,2,3,4-tetrahydro-2-methyl-3-(2-methylphenyl)-4-oxo- (9CI) (CA INDEX NAME)



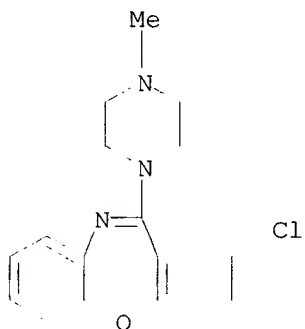
RN 27833-64-3 HCAPLUS

CN Butanedioic acid, compd. with 2-chloro-11-(4-methyl-1-piperazinyl)dibenz[b,f][1,4]oxazepine (1:1) (9CI) (CA INDEX NAME)

CM 1

CRN 1977-10-2

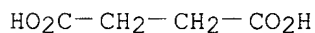
CMF C18 H18 Cl N3 O



CM 2

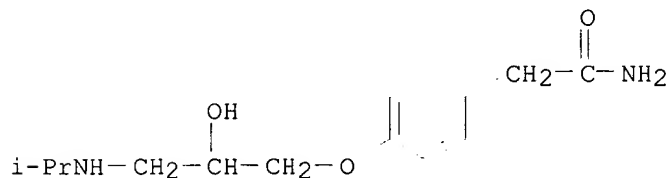
CRN 110-15-6

CMF C4 H6 O4



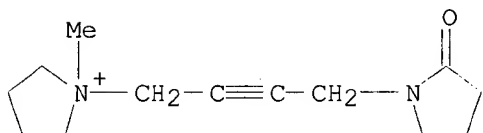
RN 29122-68-7 HCAPLUS

CN Benzeneacetamide, 4-[2-hydroxy-3-[(1-methylethyl)amino]propoxy]- (9CI) (CA INDEX NAME)



RN 30817-59-5 HCAPLUS

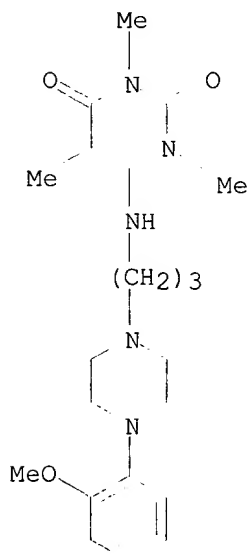
CN Pyrrolidinium, 1-methyl-1-[4-(2-oxo-1-pyrrolidinyl)-2-butynyl]-, iodide  
(8CI, 9CI) (CA INDEX NAME)



● I<sup>-</sup>

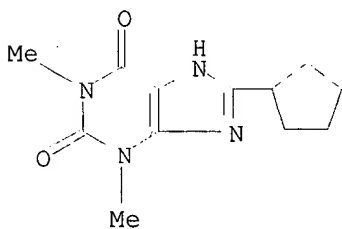
RN 34661-85-3 HCAPLUS

CN 2,4(1H,3H)-Pyrimidinedione, 6-[[3-[4-(2-methoxyphenyl)-1-piperazinyl]propyl]amino]-1,3,5-trimethyl- (9CI) (CA INDEX NAME)



RN 35873-49-5 HCAPLUS

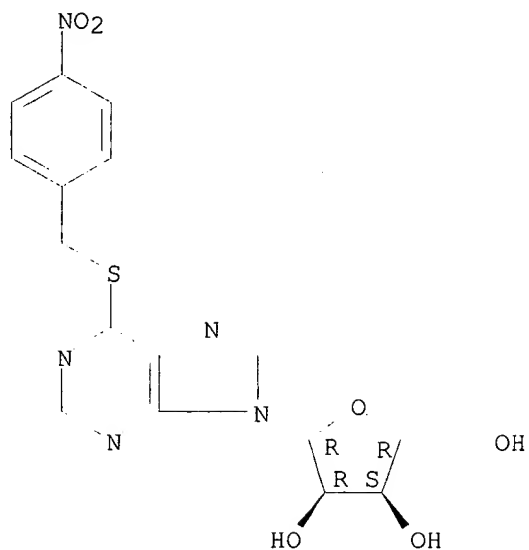
CN 1H-Purine-2,6-dione, 8-cyclopentyl-3,7-dihydro-1,3-dimethyl- (9CI) (CA INDEX NAME)



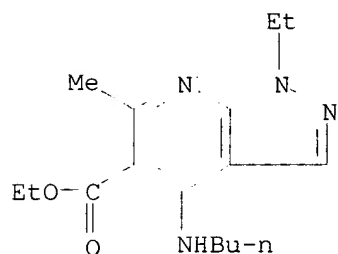
RN 38048-32-7 HCAPLUS

CN Inosine, 6-S-[(4-nitrophenyl)methyl]-6-thio- (9CI) (CA INDEX NAME)

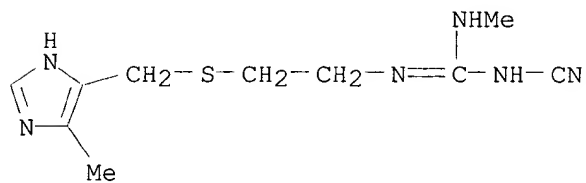
Absolute stereochemistry.



RN 41094-88-6 HCAPLUS  
 CN 1H-Pyrazolo[3,4-b]pyridine-5-carboxylic acid, 4-(butylamino)-1-ethyl-6-methyl-, ethyl ester (9CI) (CA INDEX NAME)

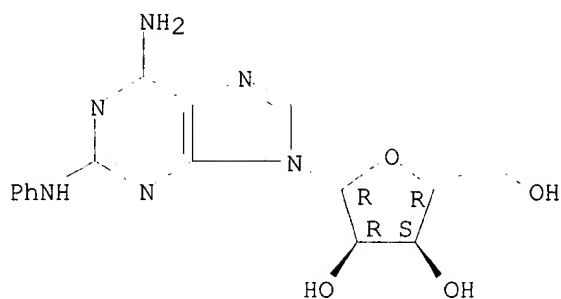


RN 51481-61-9 HCAPLUS  
 CN Guanidine, N-cyano-N'-methyl-N''-[2-[[[5-methyl-1H-imidazol-4-yl)methyl]thio]ethyl]- (9CI) (CA INDEX NAME)



RN 53296-10-9 HCAPLUS  
 CN Adenosine, 2-(phenylamino)- (9CI) (CA INDEX NAME)

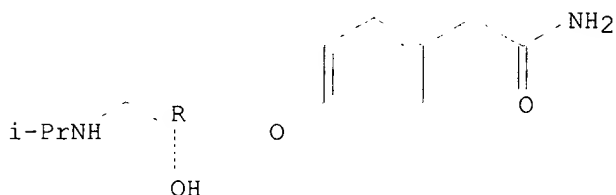
Absolute stereochemistry.



RN 56715-13-0 HCAPLUS

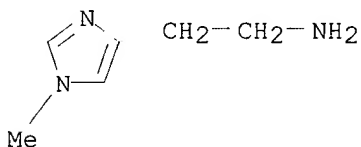
CN Benzeneacetamide, 4-[(2R)-2-hydroxy-3-[(1-methylethyl)amino]propoxy]- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).



RN 64710-63-0 HCAPLUS

CN 1H-Imidazole-4-ethanamine, 1-methyl-, monohydrochloride (9CI) (CA INDEX NAME)

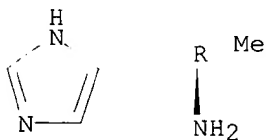


● HCl

RN 75614-89-0 HCAPLUS

CN 1H-Imidazole-4-ethanamine, .alpha.-methyl-, dihydrochloride, (.alpha.R)- (9CI) (CA INDEX NAME)

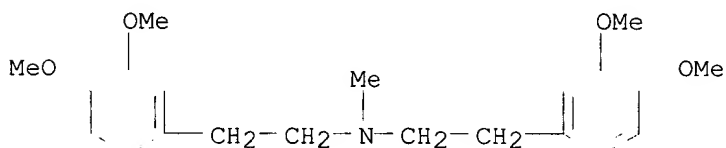
Absolute stereochemistry. Rotation (-).



●2 HCl

RN 89805-39-0 HCAPLUS

CN Benzeneethanamine, N-[2-(3,4-dimethoxyphenyl)ethyl]-3,4-dimethoxy-N-methyl-, hydrochloride (9CI) (CA INDEX NAME)

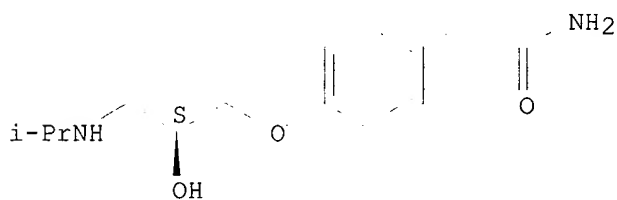


● HCl

RN 93379-54-5 HCAPLUS

CN Benzeneacetamide, 4-[(2S)-2-hydroxy-3-[(1-methylethyl)amino]propoxy]-(9CI) (CA INDEX NAME)

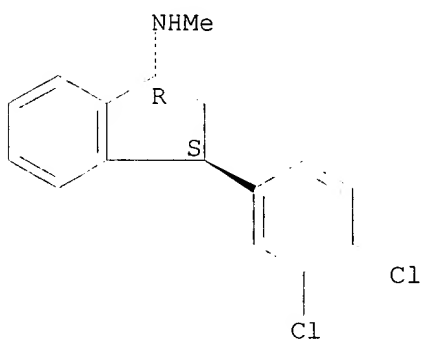
Absolute stereochemistry. Rotation (-).



RN 96850-13-4 HCAPLUS

CN 1H-Inden-1-amine, 3-(3,4-dichlorophenyl)-2,3-dihydro-N-methyl-, hydrochloride, (1R,3S)-rel- (9CI) (CA INDEX NAME)

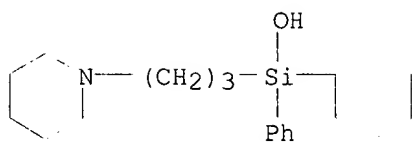
Relative stereochemistry.



● HCl

RN 100069-68-9 HCAPLUS

CN Silanol, cyclohexylphenyl[3-(1-piperidiny)propyl]-, hydrochloride (9CI) (CA INDEX NAME)

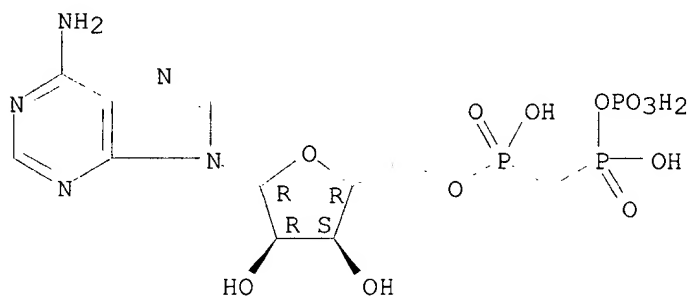


● HCl

RN 104809-20-3 HCAPLUS

CN Adenosine, 5'-[hydrogen [[hydroxy(phosphonooxy)phosphinyl]methyl]phosphonate], dilithium salt (9CI) (CA INDEX NAME)

Absolute stereochemistry.



● 2 Li

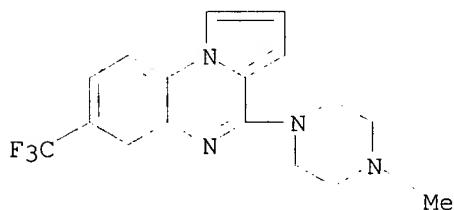
RN 109028-10-6 HCAPLUS

CN Pyrrolo[1,2-a]quinoxaline, 4-(4-methyl-1-piperazinyl)-7-(trifluoromethyl)-, (2Z)-2-butenedioate (1:2) (9CI) (CA INDEX NAME)

CM 1

CRN 109028-09-3

CMF C17 H17 F3 N4

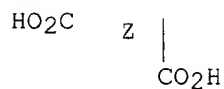


CM 2

CRN 110-16-7

CMF C4 H4 O4

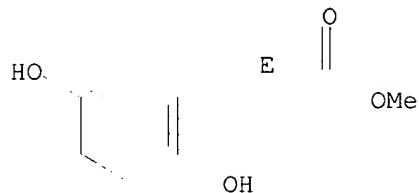
Double bond geometry as shown.



RN 123064-80-2 HCAPLUS

CN 2-Propenoic acid, 3-(2,5-dihydroxyphenyl)-, methyl ester, (2E)- (9CI) (CA INDEX NAME)

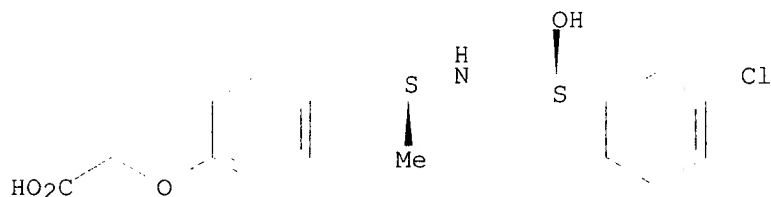
Double bond geometry as shown.



RN 127299-93-8 HCAPLUS

CN Acetic acid, [4-[(2R)-2-[[[(2R)-2-(3-chlorophenyl)-2-hydroxyethyl]amino]propyl]phenoxy]-, monosodium salt, rel- (9CI) (CA INDEX NAME)

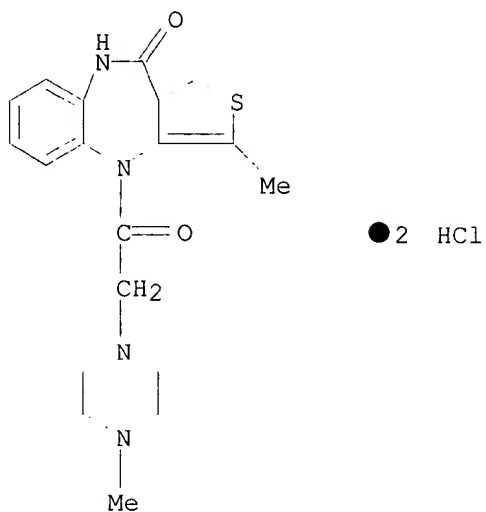
Relative stereochemistry.



● Na

RN 147416-96-4 HCAPLUS

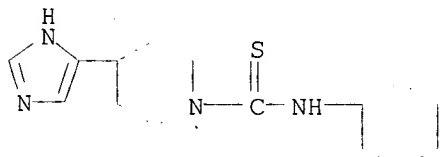
CN 10H-Thieno[3,4-b][1,5]benzodiazepin-10-one, 4,9-dihydro-3-methyl-4-[(4-methyl-1-piperazinyl)acetyl]-, dihydrochloride (9CI) (CA INDEX NAME)



RN 148440-81-7 HCAPLUS  
 CN 1-Piperidinecarbothioamide, N-cyclohexyl-4-(1H-imidazol-4-yl)-,  
 (2Z)-2-butenedioate (9CI) (CA INDEX NAME)

CM 1

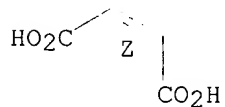
CRN 106243-16-7  
 CMF C15 H24 N4 S



CM 2

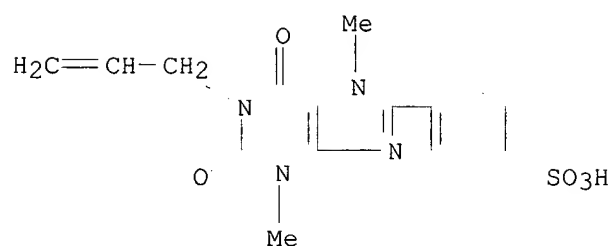
CRN 110-16-7  
 CMF C4 H4 O4

Double bond geometry as shown.



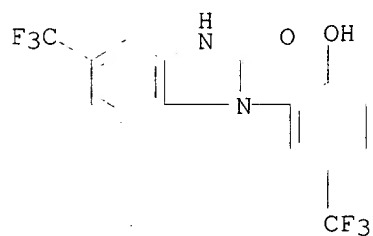
RN 149981-25-9 HCAPLUS  
 CN Benzenesulfonic acid, 4-[2,3,6,7-tetrahydro-3,7-dimethyl-2,6-dioxo-1-(2-propenyl)-1H-purin-8-yl]- (9CI) (CA INDEX NAME)





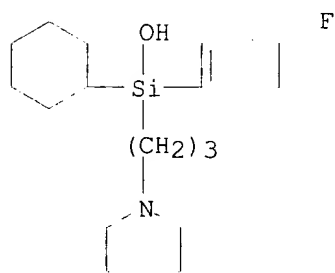
RN 153587-01-0 HCAPLUS

CN 2H-Benzimidazol-2-one, 1,3-dihydro-1-[2-hydroxy-5-(trifluoromethyl)phenyl]-5-(trifluoromethyl)- (9CI) (CA INDEX NAME)



RN 175615-76-6 HCAPLUS

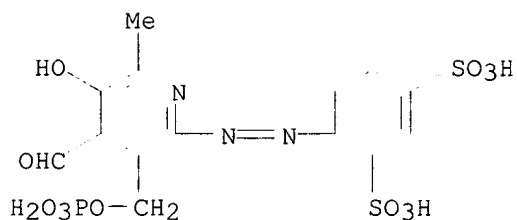
CN Silanol, cyclohexyl(4-fluorophenyl)[3-(1-piperidinyl)propyl]-, hydrochloride (9CI) (CA INDEX NAME)



● HCl

RN 192575-19-2 HCAPLUS

CN 1,3-Benzenedisulfonic acid, 4-[[4-formyl-5-hydroxy-6-methyl-3-[(phosphonoxy)methyl]-2-pyridinyl]azo]-, tetrasodium salt (9CI) (CA INDEX NAME)

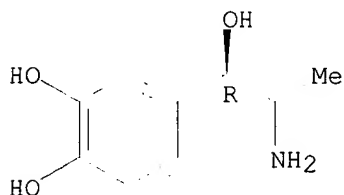


● 4 Na

RN 369647-58-5 HCAPLUS

CN 1,2-Benzenediol, 4-[(1R)-2-amino-1-hydroxypropyl]- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



RN 369647-59-6 HCAPLUS

CN 1,2-Ethanedione, 1-(4-hydroxy-3-methoxyphenyl)-, compd. with diethyldiazene (2:1) (9CI) (CA INDEX NAME)

CM 1

CRN 821-14-7

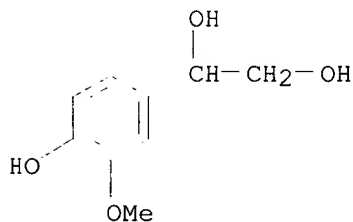
CMF C4 H10 N2

Et-N=N-Et

CM 2

CRN 534-82-7

CMF C9 H12 O4

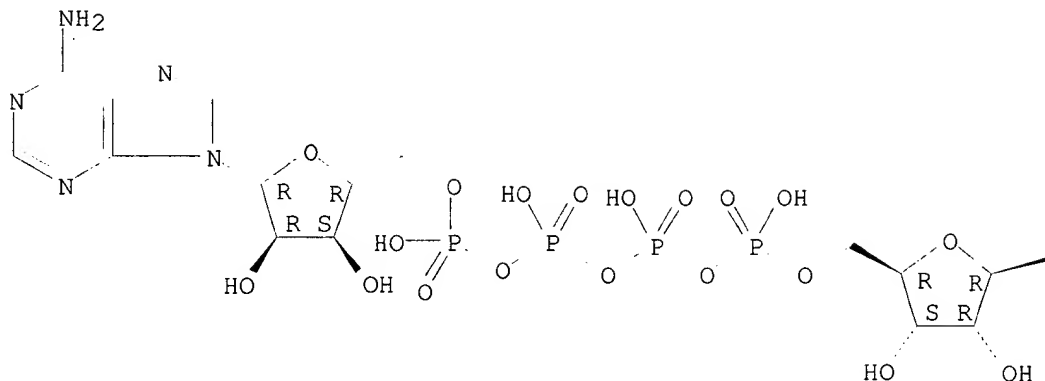


RN 369647-60-9 HCAPLUS

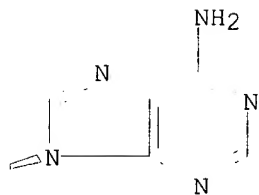
CN Adenosine 5'-(pentahydrogen tetraphosphate), P'''-fwdarw.5'-ester with adenosine, triammonium salt (9CI) (CA INDEX NAME)

Absolute stereochemistry.

PAGE 1-A

● 3 NH<sub>3</sub>

PAGE 1-B



L127 ANSWER 12 OF 28 HCAPLUS COPYRIGHT 2003 ACS

AN 2001:598462 HCAPLUS

DN 135:177709

TI Treatment and diagnosis of **Alzheimer's** disease with  
anti-Chlamydia pneumoniae agentsIN Balin, Brian J.; Abrams, J. Todd; Hudson, Alan P.; Whittum-Hudson, Judith  
A.

PA USA

SO U.S. Pat. Appl. Publ., 42 pp.

CODEN: USXXCO

DT **Patent**

LA English

IC ICM A61K031-70

ICS A01N043-04

NCL 514029000

CC 9-10 (Biochemical Methods)

Section cross-reference(s): 1, 14, 15

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	US 2001014670	A1	20010816	US 1999-227749	19990108 <--
PRAI	US 1998-70855P	P	19980109	<--	

- AB The invention relates to a method of treating **Alzheimer's** disease in a mammal comprising administering to the mammal an anti-microbial agent having anti-Chlamydia pneumoniae activity. The invention also relates to a method of diagnosing **Alzheimer's** disease in a mammal comprising measuring the serum anti-Chlamydia pneumoniae antibody titer in a patient suspected of having **Alzheimer's** disease (AD). Immunohistochem. anal. of tissues from affected regions of AD brains and congruent regions from non-AD control brains was performed to identify specific area(s) and host cell types within which the bacterium resides. Immunohistochem. anal. confirmed the presence of C. pneumoniae in affected AD brain regions and localized the bacterium to non-neuronal cells. At least three cell types, astroglia, microglia, and pericytes, were shown to harbor C. pneumoniae in the AD brain.
- ST **Alzheimer** disease treatment diagnosis Chlamydia pneumoniae; antibody Chlamydia pneumoniae blood **Alzheimer** diagnosis; antimicrobial Chlamydia **Alzheimer** disease treatment
- IT rRNA  
RL: ANT (Analyte); BOC (Biological occurrence); BSU (Biological study, unclassified); THU (Therapeutic use); ANST (Analytical study); BIOL (Biological study); OCCU (Occurrence); USES (Uses)  
(16 S, detection of microbial gene for; treatment and diagnosis of **Alzheimer's** disease with anti-Chlamydia pneumoniae agents)
- IT **Brain**  
(Chlamydia pneumoniae localization in, of humans with **Alzheimer's** disease; treatment and diagnosis of **Alzheimer's** disease with anti-Chlamydia pneumoniae agents)
- IT Apolipoproteins  
RL: BOC (Biological occurrence); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); OCCU (Occurrence); USES (Uses)  
(E, genotype allele; treatment and diagnosis of **Alzheimer's** disease with anti-Chlamydia pneumoniae agents)
- IT Test kits  
(ELISA; treatment and diagnosis of **Alzheimer's** disease with anti-Chlamydia pneumoniae agents)
- IT Glycoproteins, specific or class  
RL: ANT (Analyte); BOC (Biological occurrence); BSU (Biological study, unclassified); THU (Therapeutic use); ANST (Analytical study); BIOL (Biological study); OCCU (Occurrence); USES (Uses)  
(MOMP (major outer membrane protein), detection of microbial gene for; treatment and diagnosis of **Alzheimer's** disease with anti-Chlamydia pneumoniae agents)
- IT PCR (polymerase chain reaction)  
(RT-PCR (reverse transcription-PCR); treatment and diagnosis of **Alzheimer's** disease with anti-Chlamydia pneumoniae agents)
- IT Proteins, specific or class  
RL: ARG (Analytical reagent use); BAC (Biological activity or effector, except adverse); BPR (Biological process); BSU (Biological study, unclassified); THU (Therapeutic use); ANST (Analytical study); BIOL (Biological study); PROC (Process); USES (Uses)  
(Sezary T-cell activating factor (SAF), antibody to; treatment and diagnosis of **Alzheimer's** disease with anti-Chlamydia pneumoniae agents)
- IT Cerebrospinal fluid  
(anal. of; treatment and diagnosis of **Alzheimer's** disease with anti-Chlamydia pneumoniae agents)
- IT Macrolides  
Sulfonamides  
Tetracyclines  
RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(antibiotics; treatment and diagnosis of **Alzheimer's** disease with anti-Chlamydia pneumoniae agents)
- IT **Nervous system**

- (central, anti-SAF antibody binding to tissue of; treatment and diagnosis of **Alzheimer's** disease with anti-Chlamydia pneumoniae agents)
- IT **Nervous system**  
(central, infection; treatment and diagnosis of **Alzheimer's** disease with anti-Chlamydia pneumoniae agents)
- IT Immunoassay  
(enzyme-linked immunosorbent assay; treatment and diagnosis of **Alzheimer's** disease with anti-Chlamydia pneumoniae agents)
- IT **Brain**  
(hippocampus, Chlamydia pneumoniae detection in, of humans with **Alzheimer's** disease; treatment and diagnosis of **Alzheimer's** disease with anti-Chlamydia pneumoniae agents)
- IT Immunoassay  
(immunoelectron microscopy; treatment and diagnosis of **Alzheimer's** disease with anti-Chlamydia pneumoniae agents)
- IT Immunoassay  
(immunohistochem.; treatment and diagnosis of **Alzheimer's** disease with anti-Chlamydia pneumoniae agents)
- IT Nucleic acid hybridization  
(in situ, in situ-Pop; treatment and diagnosis of **Alzheimer's** disease with anti-Chlamydia pneumoniae agents)
- IT Nucleic acid hybridization  
(in situ; treatment and diagnosis of **Alzheimer's** disease with anti-Chlamydia pneumoniae agents)
- IT Animal cell  
Macrophage  
Monocyte  
**Oligodendrocyte**  
(infected with Chlamydia pneumoniae, for drug screening; treatment and diagnosis of **Alzheimer's** disease with anti-Chlamydia pneumoniae agents)
- IT **Astrocyte**  
(infection, with Chlamydia pneumoniae, for drug screening; treatment and diagnosis of **Alzheimer's** disease with anti-Chlamydia pneumoniae agents)
- IT Nose  
(intranasal sample of; treatment and diagnosis of **Alzheimer's** disease with anti-Chlamydia pneumoniae agents)
- IT Drug delivery systems  
(intrathecal; treatment and diagnosis of **Alzheimer's** disease with anti-Chlamydia pneumoniae agents)
- IT Antibodies  
RL: ARG (Analytical reagent use); THU (Therapeutic use); ANST (Analytical study); BIOL (Biological study); USES (Uses)  
(labeled, to Sezary T-cell activating factor (SAF); treatment and diagnosis of **Alzheimer's** disease with anti-Chlamydia pneumoniae agents)
- IT Antibiotics  
(macrolide; treatment and diagnosis of **Alzheimer's** disease with anti-Chlamydia pneumoniae agents)
- IT **Neuroglia**  
(microglia, cells infected with Chlamydia pneumoniae, for drug screening; treatment and diagnosis of **Alzheimer's** disease with anti-Chlamydia pneumoniae agents)
- IT Lipopolysaccharides  
RL: ANT (Analyte); BOC (Biological occurrence); BPR (Biological process); BSU (Biological study, unclassified); THU (Therapeutic use); ANST (Analytical study); BIOL (Biological study); OCCU (Occurrence); PROC (Process); USES (Uses)  
(monoclonal antibody to; treatment and diagnosis of **Alzheimer's** disease with anti-Chlamydia pneumoniae agents)
- IT Antibodies

RL: ARG (Analytical reagent use); THU (Therapeutic use); ANST (Analytical study); BIOL (Biological study); USES (Uses)  
 (monoclonal, to Sezary T-cell activating factor (SAF); treatment and diagnosis of **Alzheimer's** disease with anti-Chlamydia pneumoniae agents)

IT Drug delivery systems  
 (nasal; treatment and diagnosis of **Alzheimer's** disease with anti-Chlamydia pneumoniae agents)

IT **Nerve**  
 (neuron, infected with Chlamydia pneumoniae, for drug screening; treatment and diagnosis of **Alzheimer's** disease with anti-Chlamydia pneumoniae agents)

IT Anti-inflammatory agents  
 (nonsteroidal; treatment and diagnosis of **Alzheimer's** disease with anti-Chlamydia pneumoniae agents)

IT DNA  
 mRNA

RL: ANT (Analyte); BOC (Biological occurrence); BSU (Biological study, unclassified); THU (Therapeutic use); ANST (Analytical study); BIOL (Biological study); OCCU (Occurrence); USES (Uses)  
 (of Chlamydia pneumoniae localization in brain of humans with **Alzheimer's** disease; treatment and diagnosis of **Alzheimer's** disease with anti-Chlamydia pneumoniae agents)

IT **Brain**  
 (olfactory bulb, Chlamydia pneumoniae detection in, of humans with **Alzheimer's** disease; treatment and diagnosis of **Alzheimer's** disease with anti-Chlamydia pneumoniae agents)

IT Drug delivery systems  
 (oral; treatment and diagnosis of **Alzheimer's** disease with anti-Chlamydia pneumoniae agents)

IT Capillary vessel  
 (pericyte, Chlamydia pneumoniae detection in, of humans with **Alzheimer's** disease; treatment and diagnosis of **Alzheimer's** disease with anti-Chlamydia pneumoniae agents)

IT Drug delivery systems  
 (systemic; treatment and diagnosis of **Alzheimer's** disease with anti-Chlamydia pneumoniae agents)

IT **Brain**  
 (temporal cortex, Chlamydia pneumoniae detection in, of humans with **Alzheimer's** disease; treatment and diagnosis of **Alzheimer's** disease with anti-Chlamydia pneumoniae agents)

IT Antibodies  
 RL: ANT (Analyte); ARG (Analytical reagent use); BPR (Biological process); BSU (Biological study, unclassified); THU (Therapeutic use); ANST (Analytical study); BIOL (Biological study); PROC (Process); USES (Uses)  
 (to Chlamydia pneumoniae or to Sezary T-cell activating factor (SAF); treatment and diagnosis of **Alzheimer's** disease with anti-Chlamydia pneumoniae agents)

IT **Alzheimer's disease**  
 Anti-inflammatory agents

Antibiotics

Antimicrobial agents

Blood analysis

Chlamydia pneumoniae

Diagnosis

Drug screening

Electron microscopy

Mammal (Mammalia)

PCR (polymerase chain reaction)

(treatment and diagnosis of **Alzheimer's** disease with anti-Chlamydia pneumoniae agents)

IT 329900-75-6, COX-2

RL: BAC (Biological activity or effector, except adverse); BSU (Biological

study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(inhibitors; treatment and diagnosis of **Alzheimer's** disease with anti-Chlamydia pneumoniae agents)

IT 50-33-9, Phenylbutazone, biological studies 53-86-1, Indomethacin 60-54-8, Tetracycline 79-57-2, Oxytetracycline 91-22-5D, Quinoline, derivs., antibiotic compds., biological studies 114-07-8, Erythromycin 564-25-0, Doxycycline 723-46-6, Sulfamethoxazole 738-70-5, Trimethoprim 2751-09-9, Troleandomycin 5104-49-4, Flurbiprofen 10118-90-8, Minocycline 13710-19-5, Tolfenamic acid 15307-86-5, Diclofenac 15687-27-1, Ibuprofen 22071-15-4, Ketoprofen 22204-53-1, Naproxen 31793-07-4, Pirprofen 33005-95-7, Tiaprofenic acid 36322-90-4, Piroxicam 38194-50-2, Sulindac 62013-04-1, Dirithromycin 81103-11-9, Clarithromycin 82419-36-1, Ofloxacin 83905-01-5, Azithromycin 85721-33-1, Ciprofloxacin

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(treatment and diagnosis of **Alzheimer's** disease with anti-Chlamydia pneumoniae agents)

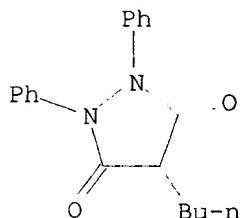
IT 50-33-9, Phenylbutazone, biological studies

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(treatment and diagnosis of **Alzheimer's** disease with anti-Chlamydia pneumoniae agents)

RN 50-33-9 HCAPLUS

CN 3,5-Pyrazolidinedione, 4-butyl-1,2-diphenyl- (6CI, 8CI, 9CI) (CA INDEX NAME)



L127 ANSWER 13 OF 28 HCAPLUS COPYRIGHT 2003 ACS

AN 2001:594375 HCAPLUS

DN 135:142289

TI Timed **pulsatile** drug delivery systems containing polymers

IN Percel, Phillip; Vishnupad, Krishna S.; Venkatesh, Gopi M.

PA Eurand America, Incorporated, USA

SO Eur. Pat. Appl., 12 pp.

CODEN: EPXXDW

DT **Patent**

LA English

IC ICM A61K009-50

ICS A61K031-18

CC 63-6 (Pharmaceuticals)

FAN.CNT 1

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
EP 1123700	A1	20010816	EP 2001-103129	20010209 <--
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO				
US 2001046964	A1	20011129	US 2001-778645	20010207 <--
WO 2001058433	A1	20010816	WO 2001-US4012	20010208 <--
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU,				

SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU,  
 ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM  
 RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, BF, BJ, CF, CG,  
 CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG

PRAI US 2000-181867P P 20000211 <--

AB A pharmaceutical dosage form such as a capsule capable of delivering therapeutic agents into the body in a time-controlled or position-controlled pulsatile release fashion, is composed of a multitude of multicoated particulates (beads, pellets, granules, etc.,) made of 1 or more populations of beads. Each of these beads except an immediate release bead has at least 2 coated membrane barriers. One of the membrane barriers is composed of an enteric polymer while the second membrane barrier is composed of a mixt. of water insol. polymer and an enteric polymer. The compn. and the thickness of the polymeric membrane barriers det. the lag time and duration of drug release from each of the bead populations. Optionally, an org. acid contg. intermediate membrane may be applied for further modifying the lag time and/or the duration of drug release. The pulsatile delivery may comprise one or more pulses to provide a plasma concn.-time profile for a therapeutic agent, predicted based on both its pharmacokinetic and pharmacodynamic considerations and in vitro/in vivo correlations. Thus, a formulation contained in the core, sotalol-HCl 8.80, sugar spheres 33.91, and povidone 0.43, in the seal coating Opadry Clear YS-1-7006 0.88, in the inner coating methacrylic acid copolymer 8.46, talc 1.69, and acetyl tri-Bu citrate 0.85, in the outer coating methacrylic acid copolymer 20.47, acetyl tri-Bu citrate 2.02, Et cellulose aq. dispersion 18.14, di-Bu sebacate 4.36% by wt., and water traces.

ST timed pulsatile drug delivery polymer

IT Monoglycerides

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (acetates; timed pulsatile drug delivery systems contg. polymers)

IT Drug delivery systems

(granules, sustained release; timed pulsatile drug delivery systems  
 contg. polymers)

IT Drug delivery systems

(pellets, sustained-release; timed pulsatile drug delivery systems  
 contg. polymers)

IT Analgesics

Anesthetics

Anti-infective agents

Anticonvulsants

**Antidiabetic agents**

**Antiparkinsonian agents**

Antirheumatic agents

Antitumor agents

Cardiovascular agents

Digestive tract

Dissolution rate

Dopamine agonists

Extrusion, nonbiological

Granulation

Milling (size reduction)

**Nervous system stimulants**

Plasticizers

Psychotropics

Spheronization

Urinary tract

(timed pulsatile drug delivery systems contg. polymers)

IT Castor oil

Shellac

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(timed pulsatile drug delivery systems contg. polymers)

IT 77-90-7, Acetyl tributyl citrate 77-93-0, Triethyl citrate 77-94-1,



Tributyl citrate 84-66-2, Diethyl phthalate 102-76-1, Triacetin 109-43-3, Dibutyl sebacate 113-45-1, Methylphenidate 114-07-8, Erythromycin 152-11-4, Verapamil hydrochloride 554-13-2, Lithium carbonate 959-24-0, Sotalol hydrochloride 9003-39-8, PVP 9004-34-6D, Cellulose, esters, biological studies 9004-57-3, Ethyl cellulose 15307-79-6, Diclofenac sodium 21829-25-4, Nifedipine 25086-15-1, Methacrylic acid-methyl methacrylate copolymer 26787-78-0, Amoxicillin 28860-95-9, Carbidopa 31677-93-7, Bupropion hydrochloride 51022-70-9, Albuterol sulfate 53237-50-6 53994-73-3, Cefaclor 56392-17-7, Metoprolol tartrate 72509-76-3, Felodipine 73590-58-6, Omeprazole 79794-75-5, Loratidine

RL: **THU (Therapeutic use)**; BIOL (Biological study); USES (Uses)  
(timed pulsatile drug delivery systems contg. polymers)

RE.CNT 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD

RE

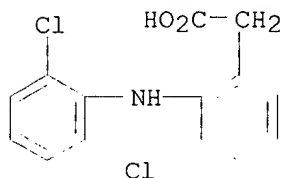
- (1) Abramowitz, R; US 5536507 A 1996 HCAPLUS
- (2) Brunetti, G; US 5900252 A 1999 HCAPLUS
- (3) Chih-Ming, C; US 5472708 A 1995 HCAPLUS
- (4) Chih-Ming, C; US 5837379 A 1998 HCAPLUS
- (5) Kinaform Technology Inc; EP 0391518 A 1990 HCAPLUS

IT 15307-79-6, Diclofenac sodium

RL: **THU (Therapeutic use)**; BIOL (Biological study); USES (Uses)  
(timed pulsatile drug delivery systems contg. polymers)

RN 15307-79-6 HCAPLUS

CN Benzeneacetic acid, 2-[(2,6-dichlorophenyl)amino]-, monosodium salt (9CI)  
(CA INDEX NAME)



● Na

L127 ANSWER 14 OF 28 HCAPLUS COPYRIGHT 2003 ACS

AN 2000:725465 HCAPLUS

DN 133:276367

TI Method for the treatment of **neurological** or **neuropsychiatric** disorders with melatonin antagonists

IN Willis, Gregory Lynn

PA Clarence Pty Ltd, Australia

SO PCT Int. Appl., 65 pp.

CODEN: PIXXD2

DT **Patent**

LA English

IC ICM A61K031-4045

ICS A61K031-165; A61K031-17; A61K031-445; A61K031-4965; A61K031-5375;  
A61K031-54; A61P025-14; A61P025-16; A61P025-18; A61P025-22;  
A61P025-24; A61P025-28

CC 1-11 (Pharmacology)

Section cross-reference(s): 14, 28

FAN.CNT 4

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2000059504	A1	20001012	WO 2000-AU275	20000331 <--

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM

RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG

US 6310085 B1 20011030 US 1999-285859 19990402 <--

BR 2000009524 A 20020219 BR 2000-9524 20000331 <--

EP 1189613 A1 20020327 EP 2000-912271 20000331 <--

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO

JP 2002541105 T2 20021203 JP 2000-609068 20000331 <--

EE 200100511 A 20021216 EE 2001-511 20000331 <--

NO 2001004674 A 20010926 NO 2001-4674 20010926 <--

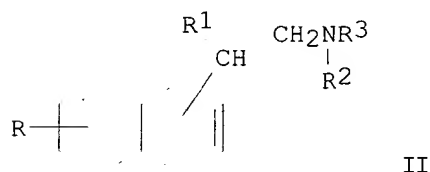
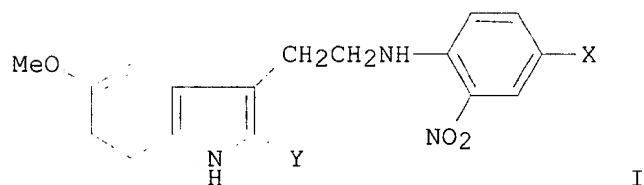
PRAI US 1999-285859 A 19990402 <--

WO 1997-AU661 A2 19971003 <--

WO 2000-AU275 W 20000331 <--

OS MARPAT 133:276367

GI



- AB A method for the treatment and/or prophylaxis of a **neurol.** or **neuropsychiatric** disorder assocd. with altered dopamine function comprises administering melatonin antagonists I (X = NO<sub>2</sub>, N<sub>3</sub>; Y = H, I) or II [R = H, OR<sub>4</sub>; R<sub>1</sub> = H, COOR<sub>5</sub>; R<sub>2</sub> = H, (substituted) alkyl; R<sub>3</sub> = CO(CH<sub>2</sub>)<sub>n</sub>R<sub>6</sub>, C(=X)NH(CH<sub>2</sub>)<sub>n</sub>R<sub>7</sub>; R<sub>4</sub> = H, (un)substituted alkyl, cycloalkyl, etc.; R<sub>5</sub> = H, (un)substituted alkyl; R<sub>6</sub> = H, (un)substituted alkyl, alkene, etc.; R<sub>7</sub> = H, (un)substituted alkyl, etc.; n = 0-3; X = O, S] to a patient in need thereof. I (X = NO<sub>2</sub>; Y = H) (ML-23) prevented the development of severe motor impairment typically exhibited by **neurotoxin** 6-hydroxydopamine-treated rats. All rats treated with ML-23 recovered and were capable of regulating their body wt.
- ST **nervous system neuropsychiatric disorder treatment**  
melatonin antagonist; movement disorder treatment ML 23; dopamine mental disorder treatment melatonin antagonist
- IT **Nervous system**  
(Huntington's chorea; **neurol.** or **neuropsychiatric** disorders treatment with melatonin antagonists)

- IT **Mental disorder**  
(Pick's disease; **neurol. or neuropsychiatric** disorders treatment with melatonin antagonists)
- IT **Mental disorder**  
(Punch drunk syndrome; **neurol. or neuropsychiatric** disorders treatment with melatonin antagonists)
- IT **Stress, animal**  
(acute; **neurol. or neuropsychiatric** disorders treatment with melatonin antagonists)
- IT **Mental disorder**  
(agoraphobia; **neurol. or neuropsychiatric** disorders treatment with melatonin antagonists)
- IT **Appetite**  
(anorexia **nervosa**; **neurol. or neuropsychiatric** disorders treatment with melatonin antagonists)
- IT **Mental disorder**  
(depression, anxiety disorders due to; **neurol. or neuropsychiatric** disorders treatment with melatonin antagonists)
- IT **Nervous system**  
(disease, malignant syndrome; **neurol. or neuropsychiatric** disorders treatment with melatonin antagonists)
- IT **Nervous system**  
(disease, multiple systems atrophy; **neurol. or neuropsychiatric** disorders treatment with melatonin antagonists)
- IT **Nervous system**  
(disease; **neurol. or neuropsychiatric** disorders treatment with melatonin antagonists)
- IT **Nervous system**  
(dystonia, acute; **neurol. or neuropsychiatric** disorders treatment with melatonin antagonists)
- IT **Nerve, disease**  
(ischemia, trans-; **neurol. or neuropsychiatric** disorders treatment with melatonin antagonists)
- IT **Behavior**  
(motor, melatonin release in relation to; **neurol. or neuropsychiatric** disorders treatment with melatonin antagonists)
- IT **Alzheimer's disease**  
Anorexia  
Anxiety  
Anxiolytics  
Cachexia  
Drug delivery systems  
    **Mental disorder**  
Movement disorders  
Multiple sclerosis  
    **Parkinson's disease**  
Phototherapy  
    **Schizophrenia**  
Veterinary medicine  
Wernicke-Korsakoff syndrome  
    (**neurol. or neuropsychiatric** disorders treatment with melatonin antagonists)
- IT **Mental disorder**  
(obsession-compulsion; **neurol. or neuropsychiatric** disorders treatment with melatonin antagonists)
- IT **Anxiety**  
(panic disorder; **neurol. or neuropsychiatric**

disorders treatment with melatonin antagonists)

IT **Mental disorder**  
(post-traumatic stress disorder; **neurol.** or **neuropsychiatric** disorders treatment with melatonin antagonists)

IT Paralysis  
(pseudobulbar; **neurol.** or **neuropsychiatric** disorders treatment with melatonin antagonists)

IT **Brain, disease**  
(stroke; **neurol.** or **neuropsychiatric** disorders treatment with melatonin antagonists)

IT **Nervous system**  
(tardive dyskinesia; **neurol.** or **neuropsychiatric** disorders treatment with melatonin antagonists)

IT 29122-68-7, Atenolol  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); **THU (Therapeutic use)**; BIOL (Biological study); USES (Uses)  
(**Parkinson's** disease treatment with bright light therapy and; **neurol.** or **neuropsychiatric** disorders treatment with melatonin antagonists)

IT 51-61-6, Dopamine, biological studies  
RL: ADV (Adverse effect, including toxicity); BIOL (Biological study) (altered; **neurol.** or **neuropsychiatric** disorders treatment with melatonin antagonists)

IT 73-31-4, Melatonin  
RL: ADV (Adverse effect, including toxicity); **THU (Therapeutic use)**; BIOL (Biological study); USES (Uses)  
(antagonists; **neurol.** or **neuropsychiatric** disorders treatment with melatonin antagonists)

IT 1199-18-4, 6-Hydroxydopamine  
RL: ADV (Adverse effect, including toxicity); BIOL (Biological study) (**neurol.** or **neuropsychiatric** disorders treatment with melatonin antagonists)

IT 152302-33-5, S-20928  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); **THU (Therapeutic use)**; BIOL (Biological study); USES (Uses)  
(**neurol.** or **neuropsychiatric** disorders treatment with melatonin antagonists)

IT 115007-18-6, ML-23  
RL: **THU (Therapeutic use)**; BIOL (Biological study); USES (Uses)  
(**neurol.** or **neuropsychiatric** disorders treatment with melatonin antagonists)

RE.CNT 9 THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD

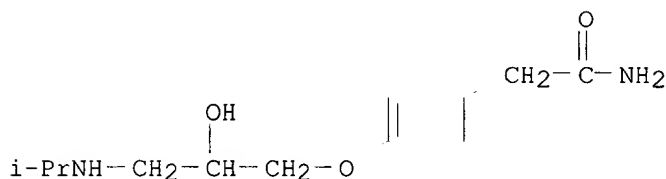
RE

(1) Adir Et Compagnie; WO 9958495 1999 HCAPLUS  
(2) Andrieux; US 5318994 1994 HCAPLUS  
(3) Lesieur; US 5385944 1995 HCAPLUS  
(4) Tenn; Brain Research 1997, V756, P293 HCAPLUS  
(5) Willis; WO 9815267 1998 HCAPLUS  
(6) Ying; European Journal of Pharmacology 1996, V296, P33 HCAPLUS  
(7) Yous; US 5420158 1995 HCAPLUS  
(8) Yous; US 5616614 1997 HCAPLUS  
(9) Zisapel; US 4880826 1989 HCAPLUS

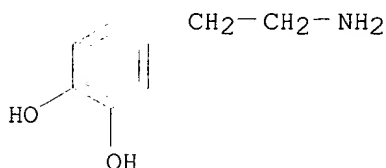
IT 29122-68-7, Atenolol  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); **THU (Therapeutic use)**; BIOL (Biological study); USES (Uses)  
(**Parkinson's** disease treatment with bright light therapy and; **neurol.** or **neuropsychiatric** disorders treatment with melatonin antagonists)

RN 29122-68-7 HCAPLUS

CN Benzeneacetamide, 4-[2-hydroxy-3-[(1-methylethyl)amino]propoxy]- (9CI)  
(CA INDEX NAME)



IT 51-61-6, Dopamine, biological studies  
RL: ADV (Adverse effect, including toxicity); BIOL (Biological study)  
(altered; **neurol.** or **neuropsychiatric** disorders  
treatment with melatonin antagonists)  
RN 51-61-6 HCAPLUS  
CN 1,2-Benzenediol, 4-(2-aminoethyl)- (9CI) (CA INDEX NAME)



L127 ANSWER 15 OF 28 HCAPLUS COPYRIGHT 2003 ACS  
AN 2000:290832 HCAPLUS  
DN 132:318003  
TI Method and composition for modulating **amyloidosis**  
IN Reiner, Peter B.; Lam, Fred Chiu-lai  
PA The University of British Columbia, Can.  
SO PCT Int. Appl., 86 pp.  
CODEN: PIXXD2  
DT **Patent**  
LA English  
IC ICM A61K031-00  
CC 1-1 (Pharmacology)  
Section cross-reference(s): 63  
FAN.CNT 3

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2000024390	A1	20000504	WO 1999-US23885	19991014 <--
W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
US 2002037843	A1	20020328	US 1998-177413	19981023 <--
US 6514686	B2	20030204		
EP 1123090	A1	20010816	EP 1999-954894	19991014 <--
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO				
JP 2002528411	T2	20020903	JP 2000-578000	19991014 <--
PRAI US 1998-177413	A2	19981023	<--	

US 1997-847616 B2 19970428 <--  
 US 1998-67523 B2 19980428 <--  
 WO 1999-US23885 W 19991014 <--

- AB Methods for modulating **amyloid** deposition in a subject are described. An effective amt. of at least one ATP-binding cassette (ABC) transporter blocker is administered to a subject, such that modulation of **amyloid** deposition occurs. Methods also include administering an effective amt. of at least one ABC transporter blocker, or a pharmaceutically acceptable salt thereof, to a subject such that a disease state assocd. with **amyloidosis** is treated. Packaged pharmaceutical compns. for treating **amyloidosis** are described. The package includes a container for holding an effective amt. of a pharmaceutical compn. and instructions for using the pharmaceutical compn. for treatment of **amyloidosis**. The pharmaceutical compn. includes at least one ABC blocker for modulating **amyloid** deposition in a subject. Methods for identifying agents which modulate **amyloid** deposition in a subject are also described. An effective amt. of at least one ATP binding cassette (ABC) transporter blocker is administered to an organism, such that modulation of **amyloid** deposition occurs.
- ST **amyloid** deposition modulation ATP binding cassette transporter blocker
- IT **Brain**  
 (ABC transporter blockade in; ATP binding cassette (ABC) transporter blocker for modulating **amyloidosis**)
- IT EST (expressed sequence tag)  
 RL: BPR (Biological process); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)  
 (ABC transporter blocker-encoding; ATP binding cassette (ABC) transporter blocker for modulating **amyloidosis**)
- IT Gene, animal  
 RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)  
 (ABC1, blockers; ATP binding cassette (ABC) transporter blocker for modulating **amyloidosis**)
- IT Gene, animal  
 RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)  
 (ABC2, blockers; ATP binding cassette (ABC) transporter blocker for modulating **amyloidosis**)
- IT Gene, animal  
 RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)  
 (ABC3, blockers; ATP binding cassette (ABC) transporter blocker for modulating **amyloidosis**)
- IT Gene, animal  
 RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)  
 (ABC7, blockers; ATP binding cassette (ABC) transporter blocker for modulating **amyloidosis**)
- IT Gene, animal  
 RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)  
 (ABC8, blockers; ATP binding cassette (ABC) transporter blocker for modulating **amyloidosis**)
- IT **Alzheimer's disease**  
**Amyloidosis**  
**Anti-Alzheimer's agents**  
 Bilayer membranes  
 Cell membrane  
 Drug screening  
 Liposomes  
 Membrane, biological

Multidrug resistance  
(ATP binding cassette (ABC) transporter blocker for modulating  
**amyloidosis**)

IT Transport proteins  
RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL  
(Biological study); PROC (Process)  
(ATP binding cassette (ABC) transporter, blockers; ATP binding cassette  
(ABC) transporter blocker for modulating **amyloidosis**)

IT Gene, animal  
RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL  
(Biological study); PROC (Process)  
(MDR1, blockers; ATP binding cassette (ABC) transporter blocker for  
modulating **amyloidosis**)

IT Gene, animal  
RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL  
(Biological study); PROC (Process)  
(MRP4, inhibitors; ATP binding cassette (ABC) transporter blocker for  
modulating **amyloidosis**)

IT Gene, animal  
RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL  
(Biological study); PROC (Process)  
(MRP5, inhibitors; ATP binding cassette (ABC) transporter blocker for  
modulating **amyloidosis**)

IT P-glycoproteins  
RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL  
(Biological study); PROC (Process)  
(blockers; ATP binding cassette (ABC) transporter blocker for  
modulating **amyloidosis**)

IT **Amyloid precursor proteins**  
RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL  
(Biological study); PROC (Process)  
(cleavage of; ATP binding cassette (ABC) transporter blocker for  
modulating **amyloidosis**)

IT **Amyloid**  
RL: ADV (Adverse effect, including toxicity); BPR (Biological process);  
BSU (Biological study, unclassified); BIOL (Biological study); PROC  
(Process)  
(deposition of; ATP binding cassette (ABC) transporter blocker for  
modulating **amyloidosis**)

IT Biological transport  
(efflux, pump; ATP binding cassette (ABC) transporter blocker for  
modulating **amyloidosis**)

IT Labels  
(for drug packaging; ATP binding cassette (ABC) transporter blocker for  
modulating **amyloidosis**)

IT Packaging materials  
(for drugs; ATP binding cassette (ABC) transporter blocker for  
modulating **amyloidosis**)

IT Head  
(injury; ATP binding cassette (ABC) transporter blocker for modulating  
**amyloidosis**)

IT Gene, animal  
RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL  
(Biological study); PROC (Process)  
(mdr3, blockers; ATP binding cassette (ABC) transporter blocker for  
modulating **amyloidosis**)

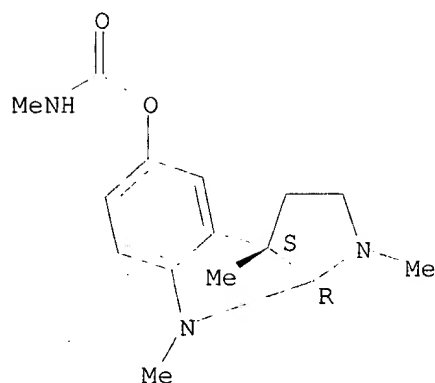
IT Blood vessel  
(microvessel, of brain, ABC transporter blockade in; ATP binding  
cassette (ABC) transporter blocker for modulating **amyloidosis**  
)

IT Protein degradation  
(of **amyloid** precursor protein; ATP binding cassette (ABC)  
transporter blocker for modulating **amyloidosis**)

- IT Transport proteins  
 RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)  
 (phospholipid-transporting, blockers; ATP binding cassette (ABC) transporter blocker for modulating **amyloidosis**)
- IT Brain, disease  
 (stroke; ATP binding cassette (ABC) transporter blocker for modulating **amyloidosis**)
- IT Amyloid  
 RL: BSU (Biological study, unclassified); MFM (Metabolic formation); BIOL (Biological study); FORM (Formation, nonpreparative)  
 (.beta.-; ATP binding cassette (ABC) transporter blocker for modulating **amyloidosis**)
- IT 50-53-3, Chlorpromazine, biological studies 50-55-5, Reserpine 51-55-8, Atropine, biological studies 52-53-9, Verapamil 54-05-7, Chloroquine 57-47-6, Physostigmine 61-54-1, Tryptamine 65-61-2, Acridine orange 83-89-6, Quinacrine 90-34-6, Primaquine 117-89-5, Trifluoperazine 130-95-0, Quinine 146-48-5, Yohimbine 260-94-6, Acridine 483-10-3, Corynanthine 485-71-2, Cinchonidine 525-66-6, Propranolol 10540-29-1, Tamoxifen 59865-13-3, Cyclosporin 59865-15-5 64657-18-7, 1,9-Dideoxyforskolin 66575-29-9, Forskolin 84371-65-3, RU-486 92302-55-1 104987-11-3, FK-506 119666-09-0, AHC-52 121584-18-7, PSC-833 129716-45-6, MS-073 140945-01-3, S9788 143664-11-3, GF120918 158681-49-3, MS-209 159997-94-1, VX-710 190454-58-1, VX-853  
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (ATP binding cassette (ABC) transporter blocker for modulating **amyloidosis**)
- IT 123955-65-7, RU 49953  
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (RU 49953; ATP binding cassette (ABC) transporter blocker for modulating **amyloidosis**)
- IT 180422-22-4, XR 9051  
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (XR 9051; ATP binding cassette (ABC) transporter blocker for modulating **amyloidosis**)
- IT 56-65-5, Atp, biological studies  
 RL: BSU (Biological study, unclassified); BIOL (Biological study)  
 (cassette binding; ATP binding cassette (ABC) transporter blocker for modulating **amyloidosis**)
- RE.CNT 1 THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD
- RE
- (1) Lam Fred Chiu Lai; WO 9848784 A 1998 HCAPLUS
- IT 57-47-6, Physostigmine  
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (ATP binding cassette (ABC) transporter blocker for modulating **amyloidosis**)
- RN 57-47-6 HCAPLUS
- CN Pyrrolo[2,3-b]indol-5-ol, 1,2,3,3a,8,8a-hexahydro-1,3a,8-trimethyl-, methylcarbamate (ester), (3aS,8aR)- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).





L127 ANSWER 16 OF 28 HCAPLUS COPYRIGHT 2003 ACS  
 AN 2000:227858 HCAPLUS  
 DN 132:260666  
 TI Identifying agents that alter mitochondrial permeability transition pores  
 and **cell death** for diagnostic and therapeutic use  
 IN Dykens, James A.; Miller, Scott W.; Ghosh, Soumitra S.; Davis, Robert E.  
 PA Mitokor, USA  
 SO PCT Int. Appl., 88 pp.  
 CODEN: PIXXD2  
 DT **Patent**  
 LA English  
 IC ICM G01N033-50  
 ICS G01N033-68; A61K031-00; C07C279-26  
 CC 1-1 (Pharmacology)  
 Section cross-reference(s): 63  
 FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2000019200	A1	20000406	WO 1999-US22261	19990924 <--
	W:	AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
	RW:	GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG			
	US 2003044776	A1	20030306	US 1998-161172	19980925 <--
	CA 2345066	AA	20000406	CA 1999-2345066	19990924 <--
	AU 9961628	A1	20000417	AU 1999-61628	19990924 <--
	EP 1116027	A1	20010718	EP 1999-948458	19990924 <--
	R:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO			
	JP 2002525630	T2	20020813	JP 2000-572655	19990924 <--
PRAI	US 1998-161172	A	19980925 <--		
	WO 1999-US22261	W	19990924 <--		

AB Methods are provided for identifying agents that affect mitochondrial functions and cell death. Such agents are useful for treating diseases assocd. with mitochondrial dysfunction and in methods of identifying a risk or presence of such diseases. In particular, the invention relates to the loss of mitochondrial membrane potential (.DELTA..PSI.m) during mitochondrial permeability transition (MPT) and further provides a measurable rate loss function, changes in which are useful e.g. for detecting agents that affect one or more mitochondrial functions, for

- detecting mitochondrial diseases, and for studying mol. components of mitochondria that regulate MPT.
- ST mitochondria permeability transition pore therapeutic identification; diagnosis mitochondrial disease permeability transition pore; cell death mitochondrial permeability therapeutic identification; membrane potential mitochondria diagnostic therapeutic identification
- IT Transport proteins  
RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)  
(ADP/ATP carrier; identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT Cyclophilins  
RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)  
(D; identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT Apolipoproteins  
RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)  
(E, genotype; identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT **Nervous system**  
(Huntington's chorea; identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT **Brain, disease**  
(MELAS (mitochondrial myopathy, encephalopathy, lactic acidosis, and stroke-like episodes); identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT Muscle, disease  
(MERRF (myoclonic epilepsy assocd. with ragged-red muscle fibers); identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT Animal cell line  
(SH-SY5Y, cybrid cell produced with; identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT Annexins  
RL: BPR (Biological process); BSU (Biological study, unclassified); BUU (Biological use, unclassified); BIOL (Biological study); PROC (Process); USES (Uses)  
(V, FITC conjugates; identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT Anion channel  
RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)  
(VDAC (voltage-dependent anion-selective channel); identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT **Neurotransmitters**  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study)  
(amino acid; identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT **Diabetes mellitus**  
(and mitochondrial diabetes and deafness; identification of agents that

- alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT Gene, animal  
RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)  
(bcl-2, Bcl-2 gene family-encoded polypeptide; identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT Membrane potential  
(biol.; identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT Transport proteins  
RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)  
(calcium-transporting, mitochondrial calcium uniporter; identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT Platelet (blood)  
(cybrid cell produced with; identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT Animal cell  
(cybrid cell; identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT Muscle, disease  
(degeneration; identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT Mitochondria  
(diseases; identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT **Nervous system**  
(dystonia; identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT Pathogen  
(eukaryotic; identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT Indicators  
(for inner mitochondrial membrane potential; identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT Eye, disease  
(hereditary optic atrophy; identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT Cell proliferation  
(hyperproliferative disease; identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT Affinity labeling  
Alzheimer's disease  
Anti-Alzheimer's agents  
Antidiabetic agents  
Antiparkinsonian agents  
Antipsychotics  
Antitumor agents  
Apoptosis  
Brain, disease  
Cell death

Cytotoxic agents  
 Diagnosis  
 Drug delivery systems  
 Drug screening  
 Electron transport system, biological  
 Fluorometry  
 Genotypes  
 Insect (Insecta)  
 Ionophores  
 Lepidoptera  
 Mitochondria  
 Necrosis  
 Neoplasm  
 Nucleic acid library  
**Parkinson's disease**  
 Plant (Embryophyta)  
 Psoriasis

**Schizophrenia**

(identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)

- IT DNA
  - RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)
  - (identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT Mitochondria
  - Mitochondria
  - (inner membrane; identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT Membrane, biological
  - Membrane, biological
  - (inner mitochondrial; identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT Biological transport
  - (intracellular, phosphatidylserine; identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT Acidosis
  - (lactic; identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT Time-of-flight mass spectrometry
  - Time-of-flight mass spectrometry
  - (laser-induced photodesorption; identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT Deafness
  - (mitochondrial diabetes and deafness; identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT Amino acids, biological studies
  - RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study)
  - (**neurotransmitter**; identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT Parasite
  - (of human; identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT Eukaryote (Eukaryotae)

- (pathogen; identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT Benzodiazepine receptors  
RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)  
(peripheral; identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT Biological transport  
(permeation; identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT Laser ionization mass spectrometry  
Laser ionization mass spectrometry  
(photodesorption, matrix-assisted; identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT Laser desorption mass spectrometry  
Laser desorption mass spectrometry  
(photoionization, matrix-assisted; identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT Proliferation inhibition  
(proliferation inhibitors; identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT Laser desorption mass spectrometry  
Laser desorption mass spectrometry  
(time-of-flight; identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT Antibodies  
RL: BPR (Biological process); BSU (Biological study, unclassified); BUU (Biological use, unclassified); BIOL (Biological study); PROC (Process); USES (Uses)  
(to cytochrome c; identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT Phosphatidylserines  
RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)  
(translocation; identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT 145037-81-6, Rhod 2  
RL: BPR (Biological process); BSU (Biological study, unclassified); BUU (Biological use, unclassified); BIOL (Biological study); PROC (Process); USES (Uses)  
(Rhod 2; identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT 9007-43-6, Cytochrome c, biological studies 122191-40-6, Caspase 1  
169592-56-7, Caspase 3 186322-81-6, Caspase  
RL: BAC (Biological activity or effector, except adverse); BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)  
(identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)
- IT 51-83-2, Carbachol 56-86-0, L-Glutamic acid, biological studies  
6384-92-5, N-Methyl-D-aspartic acid 11076-19-0, Bongkreikic acid  
11103-72-3, Ruthenium red 17754-44-8, Atractyloside 56092-81-0, Ionomycin  
67526-95-8, Thapsigargin 79217-60-0, Cyclosporin  
169332-61-0 182374-54-5D, derivs. 201608-13-1 217174-04-4

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study)  
 (identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)

IT 102-02-3, 1-Phenylbiguanide  
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); **THU (Therapeutic use)**; BIOL (Biological study); USES (Uses)  
 (identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)

IT 7440-70-2, Calcium, biological studies  
 RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)  
 (identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)

IT 2156-29-8 3520-43-2, JC-1 18198-39-5, Tetraphenylphosphonium 27072-45-3D, Fluorescein isothiocyanate, annexin V conjugates 30827-04-4, Rhodamine B hexyl ester 53213-82-4, DiOC6(3) 62669-70-9, Rhodamine 123 115532-49-5 137993-41-0, Rhodamine 800 139626-15-6, Tetramethylrhodamine ethyl ester  
 RL: BPR (Biological process); BSU (Biological study, unclassified); BUU (Biological use, unclassified); BIOL (Biological study); PROC (Process); USES (Uses)  
 (identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)

IT 9001-15-4, Creatine kinase  
 RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)  
 (mitochondrial intermembrane; identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)

IT 9001-51-8, Hexokinase  
 RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)  
 (mitochondrial-assocd.; identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)

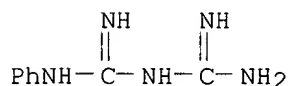
RE.CNT 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD

RE  
 (1) Beal, M; Biochimica et Biophysica Acta 1998, V1366(1-2), P211 HCAPLUS  
 (2) Diamond, J; GB 1410925 A 1975 HCAPLUS  
 (3) Friberg, H; Journal of Neuroscience 1998, V18(14), P5151 HCAPLUS  
 (4) Hirsch, T; Cell Biology and Toxicology 1998, V4(2), P141

IT 102-02-3, 1-Phenylbiguanide  
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); **THU (Therapeutic use)**; BIOL (Biological study); USES (Uses)  
 (identification of agents that alter mitochondrial permeability transition pores and cell death for diagnostic and therapeutic use)

RN 102-02-3 HCAPLUS

CN Imidodicarbonimidic diamide, N-phenyl- (9CI) (CA INDEX NAME)



L127 ANSWER 17 OF 28 HCAPLUS COPYRIGHT 2003 ACS

AN 2000:169319 HCAPLUS

DN 132:212709

TI Pharmaceutical composition containing tarcine for the treatment of

**neurological diseases**

IN Guittard, George V.; Childers, Jerry D.; Wong, Patrick S. L.; Gumucio, Fernando E.; Kidney, David J.

PA Alza Corporation, USA

SO U.S., 16 pp., Cont.-in-part of U.S. 5,698,224.

CODEN: USXXAM

DT **Patent**

LA English

IC ICM A61K009-00

ICS A61K031-13; A61K031-135

NCL 424457000

CC 63-6 (Pharmaceuticals)

Section cross-reference(s): 1

FAN.CNT 2

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	US 6036973	A	20000314	US 1997-892995	19970715 <--
	US 5698224	A	19971216	US 1994-266045	19940627 <--
	CA 2187332	AA	19960104	CA 1995-2187332	19950614 <--
PRAI	US 1994-266045		19940627	<--	

AB A dosage form is disclosed for administering 10 ng to 1200 mg tacrine to a patient in need of tacrine therapy. A core comprising 86.15 mg of tacrine hydrochloride, 86.15 mg of mannitol, 7.25 mg of poly(vinylpyrrolidone) and 1.81 mg of magnesium stearate was prepd. A semipermeable wall was coated around the individual, sep. cores comprising 80 % cellulose acetate having a 39.8% acetyl content and 20 % poly(vinylpyrrolidone). An exit passageway was drilled through the semipermeable wall connecting the tacrine with the exterior of each dosage form. The exit port had a diam. of 30 mils (0.76 mm) and each dosage form dispensed tacrine for 24 h.

ST pharmaceutical tablet tacrine **neurol** disease

IT **Nervous system**

(disease; pharmaceutical compn. contg. tacrine for treatment of **neurol.** diseases)

IT **Alzheimer's disease**

(pharmaceutical compn. contg. tacrine for treatment of **neurol** . diseases)

IT Estrogens

Phosphatidylserines

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(pharmaceutical compn. contg. tacrine for treatment of **neurol** . diseases)

IT Polyoxyalkylenes, biological studies

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(pharmaceutical compn. contg. tacrine for treatment of **neurol** . diseases)

IT Drug delivery systems

(sustained-release; pharmaceutical compn. contg. tacrine for treatment of **neurol.** diseases)

IT **57-47-6**, Physostigmine 59-02-9, .alpha.-Tocopherol 321-64-2, Tacrine 485-35-8, Cytisine **504-24-5**, Fampridine 1684-40-8, Tacrine hydrochloride 14611-51-9, Selegiline 26445-05-6, Aminopyridine 55242-55-2, Propentofylline 66085-59-4, Nimodipine 68497-62-1, Pramiracetam 72432-10-1, ANiracetam 90293-01-9, Bifemelane 105431-72-9, Linopirdine 120014-06-4, Donepezil 124027-47-0, 1-Hydroxy-tacrine 174528-42-8, Tacrine hydrobromide 174528-43-9, Tacrine sulfate 174528-44-0, Tacrine phosphate 174528-45-1, Tacrine lactate 174528-46-2, Tacrine citrate 174528-47-3, Tacrine malate 174528-48-4, Tacrine maleate 174528-49-5, Tacrine fumarate 174528-50-8 174528-51-9 174528-52-0, Tacrine aspartate 174528-53-1, Tacrine salicylate 174528-54-2, Tacrine edisylate 174528-55-3, Tacrine laurate 174528-56-4, Tacrine palmitate 174528-57-5, Tacrine nitrate

174528-58-6, Tacrine borate 174528-59-7, Tacrine acetate 174528-60-0, Tacrine oleate 174672-18-5, Tacrine tartrate  
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); **THU (Therapeutic use)**; BIOL (Biological study); USES (Uses)

(pharmaceutical compn. contg. tacrine for treatment of **neuro**l . diseases)

IT 50-70-4, Sorbitol, biological studies 9003-39-8, Poly(vinylpyrrolidone) 9004-32-4, Sodium CM-cellulose 9004-35-7, Cellulose acetate 9004-64-2, Hydroxypropylcellulose 9004-65-3, Hydroxypropylmethylcellulose 25322-68-3, Polyethylene glycol;

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (pharmaceutical compn. contg. tacrine for treatment of **neuro**l . diseases)

RE.CNT 20 THERE ARE 20 CITED REFERENCES AVAILABLE FOR THIS RECORD  
 RE

- (1) Anon; WO 9215285 1992 HCAPLUS
- (2) Anon; WO 9324154 1993 HCAPLUS
- (3) Anon; EP 0595365 A1 1994 HCAPLUS
- (4) Anon; WO 9503052 1995 HCAPLUS
- (5) Cortese; US 4327725 1982
- (6) Guittard; US 5698224 1997 HCAPLUS
- (7) Higuchi; US 5916925 1999 HCAPLUS
- (8) Saunders; US 4063064 1977
- (9) Stephen; US 4857330 1989
- (10) Summers; US 4816456 1989 HCAPLUS
- (11) Theeuwes; US 3845770 1974 HCAPLUS
- (12) Theeuwes; US 3916899 1975 HCAPLUS
- (13) Theeuwes; US 4077407 1978 HCAPLUS
- (14) Theeuwes; US 4088864 1978
- (15) Wong; US 4612008 1986
- (16) Wong; US 4765989 1988 HCAPLUS
- (17) Wong; US 4783337 1988 HCAPLUS
- (18) Wurster; US 2799241 1957
- (19) Wurster, D; J Am Phar Assoc, Sci Ed 1959, V48, P451 MEDLINE
- (20) Wurster, D; J Am Phar Assoc, Sci Ed 1960, V49, P82 MEDLINE

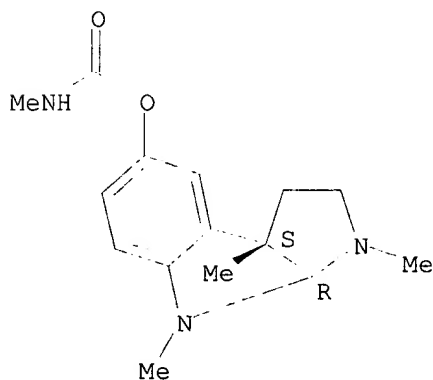
IT 57-47-6, Physostigmine 504-24-5, Fampridine  
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); **THU (Therapeutic use)**; BIOL (Biological study); USES (Uses)

(pharmaceutical compn. contg. tacrine for treatment of **neuro**l . diseases)

RN 57-47-6 HCAPLUS

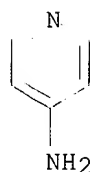
CN Pyrrolo[2,3-b]indol-5-ol, 1,2,3,3a,8,8a-hexahydro-1,3a,8-trimethyl-, methylcarbamate (ester), (3aS,8aR)- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).





RN 504-24-5 HCAPLUS  
 CN 4-Pyridinamine (9CI) (CA INDEX NAME)



L127 ANSWER 18 OF 28 HCAPLUS COPYRIGHT 2003 ACS

AN 1999:672562 HCAPLUS

DN 131:281590

TI Methods for treating **neuropsychiatric** disorders

IN Tsai, Guochuan; Coyle, Joseph

PA The General Hospital Corporation, USA

SO PCT Int. Appl., 27 pp.

CODEN: PIXXD2

DT **Patent**

LA English

IC ICM A61K031-00

CC 1-11 (Pharmacology)

Section cross-reference(s): 63

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 9952519	A2	19991021	WO 1999-US8056	19990414 <--
	WO 9952519	A3	19991202		
	W:	AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
	RW:	GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG			
	CA 2328197	AA	19991021	CA 1999-2328197	19990414 <--
	AU 9935571	A1	19991101	AU 1999-35571	19990414 <--
	EP 1073432	A2	20010207	EP 1999-917453	19990414 <--
	R:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI			
	US 6228875	B1	20010508	US 1999-291296	19990414 <--
	JP 2002511409	T2	20020416	JP 2000-543129	19990414 <--
	US 2002035145	A1	20020321	US 2001-834351	20010413 <--
	US 6420351	B1	20020716		
	US 2002193429	A1	20021219	US 2002-196686	20020715 <--
PRAI	US 1998-81645P	P	19980414	<--	
	US 1999-291296	A1	19990414	<--	
	WO 1999-US8056	W	19990414	<--	
	US 2001-834351	A1	20010413		
AB	The invention provides methods for treating <b>neuropsychiatric</b> disorders such as schizophrenia, <b>Alzheimer's</b> Disease, autism, depression, benign forgetfulness, childhood learning disorders, close head injury, and attention deficit disorder. The methods entail administering to a patient with a <b>neuropsychiatric</b> disorder a pharmaceutical compn. contg. (i) a therapeutically effective amt. of D-alanine (or a modified form), provided that the compn. is substantially free of D-cycloserine, and/or (ii) D-serine (or a modified form), and/or (iii) 105 to 500 mg of D-cycloserine (or a modified form), and/or (iv)				

N-methylglycine (or a modified form). Using double-blind conditions, patients were randomly assigned to receive placebo (fruit juice), D-serine 30, D-alanine 60-100, or N-methylglycine 30 mg/kg/day once a day by mouth for 6 wk. Treatment with D-serine, D-alanine, or N-methylglycine improved the schizophrenic symptoms and **cognitive** deficit of the patients. Specifically, treatment with D-serine resulted in a 21% redn. of the neg. symptoms (on the SANS scale), and it resulted in a 17% redn. of the pos. symptoms. Treatment with D-alanine resulted in an 11% redn. of the neg. symptoms and a 12% redn. of the pos. symptoms. Reatment with N-methylglycine resulted in a 20% redn. of the neg. symptoms and a 15% redn. of the pos. symptoms. These redns. in the neg. and pos. symptoms represented clin. significant improvement.

ST **neuropsychiatric** disorder treatment serine alanine cycloserine;  
methylglycine **neuropsychiatric** disorder treatment;  
antidepressant serine alanine cycloserine

IT **Mental disorder**  
(attention deficit disorder; methods for treating  
**neuropsychiatric** disorders)

IT **Mental disorder**  
(autism; methods for treating **neuropsychiatric** disorders)

IT **Anti-Alzheimer's agents**

Antidepressants

Antipsychotics

**Cognition enhancers**

**Mental disorder**

Psychostimulants

**Schizophrenia**

(methods for treating **neuropsychiatric** disorders)

IT 50-47-5, Desipramine 50-48-6, Amitriptyline 50-49-7, Imipramine  
50-52-2, Thioridazine 50-53-3, Chlorpromazine, biological studies  
51-64-9, Dextroamphetamine 51-71-8, Phenelzine 52-86-8, Haloperidol  
58-39-9, Perphenazine 68-41-7, D-CycloSerine 68-41-7D, D-CycloSerine,  
derivs. 68-41-7D, D-CycloSerine, salts and complexes 69-23-8,  
Fluphenazine 72-69-5, Nortriptyline 107-43-7, N,N,N-TriMethylglycine  
107-97-1, N-Methylglycine 107-97-1D, N-Methylglycine, derivs.  
113-45-1, Methylphenidate 113-59-7, Chlorprothixene 117-89-5,  
Trifluoperazine 155-09-9, TRanylcypromine 303-49-1, Clomipramine  
312-84-5, D-Serine 312-84-5D, D-Serine, derivs. 321-64-2, Tacrine  
338-69-2, D-Alanine 338-69-2D, D-Alanine, derivs. 438-60-8,  
Protriptyline 537-46-2, Methamphetamine 548-73-2, Droperidol  
653-03-2, Butaperazine 1118-68-9, N,N,-DiMethylglycine 1668-19-5,  
Doxepin 1977-10-2, Loxapine 2062-78-4, Pimozide 2152-34-3, Pemoline  
2622-30-2, Carphenazine 2746-81-8, Fluphenazine enanthate 2751-68-0,  
Acetophenazine 3313-26-6, Thiothixene 3819-00-9, Piperacetazine  
5002-47-1, Fluphenazine decanoate 5588-33-0, Mesoridazine 5786-21-0,  
Clozapine 7416-34-4, Molindone 10262-69-8, Maprotiline 14028-44-5,  
Amoxapine 15676-16-1, Sulpiride 15975-28-7 19794-93-5,  
Trazodone 32342-58-8, Sodium D-Alanine 34911-55-2, Bupropion  
54739-18-3, Fluvoxamine 54910-89-3, Fluoxetine 61869-08-7  
74050-97-8, Haloperidol decanoate 79617-96-2, Sertraline 80125-14-0,  
Remoxipride 83366-66-9, Nefazodone 85650-52-8, Mirtazapine  
106266-06-2, Risperidone 109026-02-0, D-Alanine, monopotassium salt  
111974-69-7, Quetiapine 120014-06-4, Donepezil 132539-06-1, Olanzapine  
146939-27-7, Ziprasidone 152005-29-3 246855-94-7 246855-95-8  
246855-97-0 246855-99-2 246875-51-4 246875-52-5 246875-53-6  
246875-54-7

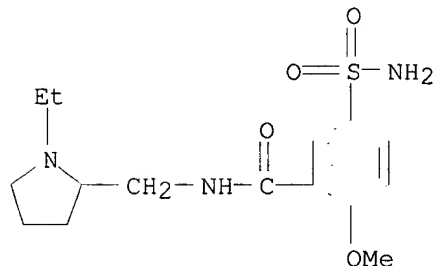
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); **THU (Therapeutic use)**; BIOL (Biological study); USES (Uses)

(methods for treating **neuropsychiatric** disorders)

IT 56-40-6, Glycine, biological studies

RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(uptake inhibitors; methods for treating **neuropsychiatric**

disorders)  
 IT 15676-16-1, Sulpiride  
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (methods for treating neuropsychiatric disorders)  
 RN 15676-16-1 HCAPLUS  
 CN Benzamide, 5-(aminosulfonyl)-N-[(1-ethyl-2-pyrrolidinyl)methyl]-2-methoxy- (9CI) (CA INDEX NAME).



L127 ANSWER 19 OF 28 HCAPLUS COPYRIGHT 2003 ACS

AN 1999:344848 HCAPLUS

DN 131:714

TI Therapeutic uses of triazolo-pyridazine derivatives

IN Castro Pineiro, Jose Luis; Hefti, Franz Fridolin; Hill, Raymond George; McKernan, Ruth; Tattersall, Frederick David; Whiting, Paul John

PA Merck Sharp & Dohme Limited, UK

SO PCT Int. Appl., 71 pp.

CODEN: PIXXD2

DT Patent

LA English

IC ICM A61K031-50

ICS A61K031-00; A61K045-06

CC 1-11 (Pharmacology)

Section cross-reference(s): 63

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 9925353	A1	19990527	WO 1998-GB3328	19981106 <--
	W:			AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM	
	RW:			GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG	
	AU 9910415	A1	19990607	AU 1999-10415	19981106 <--
	US 6174886	B1	20010116	US 1998-191304	19981112 <--
	US 6107296	A	20000822	US 1998-206416	19981207 <--
	US 6110915	A	20000829	US 1998-208288	19981208 <--
	US 6046196	A	20000404	US 1998-208291	19981209 <--
	US 6063783	A	20000516	US 1998-209071	19981210 <--
PRAI	GB 1997-23999	A	19971113	<--	
	GB 1997-26699	A	19971218	<--	
	GB 1997-26700	A	19971218	<--	
	GB 1997-26701	A	19971218	<--	
	GB 1997-26702	A	19971218	<--	
	GB 1998-1581	A	19980123	<--	

WO 1998-GB3328 W 19981106 <--  
OS MARPAT 131:714  
AB A class of substituted or 7,8-ring fused 1,2,4-triazolo[4,3-b]pyridazine derivs., possessing an optionally substituted cycloalkyl, Ph or heteroaryl substituent at the 3-position and a substituted alkoxy moiety at the 6-position, are selective ligands for GABAA receptors, in particular having high affinity for the .alpha.2 and/or .alpha.3 subunit thereof, and are accordingly of benefit in the treatment and/or prevention of psychotic disorders including schizophrenia; **neurodegeneration** arising from cerebral ischemia; pain; emesis; and muscle spasm or spasticity, e.g. in paraplegic patients.  
ST triazolopyridazine deriv GABAA ligand therapeutic; antipsychotic schizophrenia analgesic antiemetic triazolopyridazine deriv; **neurodegeneration** cerebral ischemia triazolopyridazine deriv; muscle spasm spasticity triazolopyridazine deriv  
IT 5-HT antagonists  
(5-HT3; triazolo-pyridazine deriv. GABAA ligands and therapeutic use, alone or with other compds.)  
IT GABA agonists  
(GABAA; triazolo-pyridazine deriv. GABAA ligands and therapeutic use, alone or with other compds.)  
IT GABA receptors  
RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)  
(GABAA; triazolo-pyridazine deriv. GABAA ligands, and therapeutic use)  
IT Tachykinin receptors  
(NK1 antagonists; triazolo-pyridazine deriv. GABAA ligands and therapeutic use, alone or with other compds.)  
IT Glutamate receptors  
RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)  
(NMDA-binding, strychnine-insensitive glycine modulatory site of NMDA receptor; triazolo-pyridazine deriv. GABAA ligands and therapeutic use, alone or with other compds.)  
IT Opioids  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(analgesic; triazolo-pyridazine deriv. GABAA ligands and therapeutic use, alone or with other compds.)  
IT **Nerve**  
(**degeneration**, from cerebral **ischemia**;  
triazolo-pyridazine deriv. GABAA ligands, and therapeutic use)  
IT **Neurotransmission**  
(glutamatergic, modulators; triazolo-pyridazine deriv. GABAA ligands and therapeutic use, alone or with other compds.)  
IT **Brain, disease**  
(**ischemia**, **neurodegeneration** from;  
triazolo-pyridazine deriv. GABAA ligands, and therapeutic use)  
IT Cytoprotective agents  
(**neuroprotectants**; triazolo-pyridazine deriv. GABAA ligands, and therapeutic use)  
IT Anti-inflammatory agents  
(nonsteroidal; triazolo-pyridazine deriv. GABAA ligands and therapeutic use, alone or with other compds.)  
IT Drug delivery systems  
(prodrugs; triazolo-pyridazine deriv. GABAA ligands, and therapeutic use)  
IT Muscle relaxants  
(spasmolytics; triazolo-pyridazine deriv. GABAA ligands, and therapeutic use)  
IT Cholinergic antagonists  
Dopamine antagonists

(triazolo-pyridazine deriv. GABAA ligands and therapeutic use, alone or with other compds.)

IT Analgesics  
Antiemetics  
Antipsychotics  
Drug delivery systems  
Muscle relaxants  
(triazolo-pyridazine deriv. GABAA ligands, and therapeutic use)

IT 39391-18-9  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(2, inhibitors; triazolo-pyridazine deriv. GABAA ligands and therapeutic use, alone or with other compds.)

IT 12794-10-4D, Benzodiazepine, derivs.  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(GABAA receptor benzodiazepine binding site; triazolo-pyridazine deriv. GABAA ligands, and therapeutic use)

IT 56-40-6, Glycine, biological studies 57-24-9, Strychnine  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(strychnine-insensitive glycine modulatory site of NMDA receptor; triazolo-pyridazine deriv. GABAA ligands and therapeutic use, alone or with other compds.)

IT 50-52-2, Thioridazine 50-53-3, Chlorpromazine, biological studies  
52-86-8, Haloperidol 58-39-9, Perphenazine 69-23-8, Fluphenazine  
113-59-7, Chlorprothixene 117-89-5, Trifluoperazine 1134-47-0,  
Baclofen 1977-10-2, Loxapine 2062-78-4, Pimozide 2751-68-0,  
Acetophenazine 3313-26-6, Thiethixene 5588-33-0, Mesoridazine  
5786-21-0, Clozapine 7416-34-4, Molindone **15676-16-1**,  
Sulpiride 71125-38-7, Meloxicam 106266-06-2, Risperidone  
127625-29-0, Fananserin 131986-45-3, Xanomeline 132539-06-1,  
Olanzapine 146939-27-7, Ziprasidone 162011-90-7, Rofecoxib  
169590-42-5, Celecoxib  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); **THU (Therapeutic use)**; BIOL (Biological study); USES (Uses)  
(triazolo-pyridazine deriv. GABAA ligands and therapeutic use, alone or with other compds.)

IT 202929-19-9 202929-20-2 202929-21-3 202929-22-4 202929-23-5  
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202930-44-7 202930-45-8 202930-46-9 202930-47-0 202930-48-1  
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202930-71-0	202930-76-5	202930-80-1	202930-83-4	202930-86-7
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225642-08-0	225642-09-1	225642-10-4	225642-11-5	225642-16-0
225642-18-2	225642-21-7	225642-28-4		

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(triazolo-pyridazine deriv. GABAA ligands, and therapeutic use)

RE.CNT 13 THERE ARE 13 CITED REFERENCES AVAILABLE FOR THIS RECORD

RE

- (1) Anon; S-TRIAZOLO(3,4-a) (5, 6, 7, 8)TETRAHYDROPHTHA LAZINES 1978, 5, HCAPLUS
- (2) Delini-Stula, A; JOURNAL OF PSYCHIATRIC RESEARCH 1996, V30(4), P239 MEDLINE
- (3) Dunn, E; SOCIETY FOR NEUROSCIENCE ABSTRACTS 1995, V21(1-3), P2046
- (4) Hadingham, K; MOLECULAR PHARMACOLOGY 1993, V43, P970 HCAPLUS
- (5) Hall, E; JOURNAL OF CEREBRAL BLOOD FLOW AND METABOLISM 1997, V17(8), P875 HCAPLUS
- (6) Knoll Ag; WO 9632393 A 1996 HCAPLUS
- (7) Lepetit Spa; EP 0085840 A 1983 HCAPLUS
- (8) Merck Sharp & Dohme; WO 9834923 A 1998 HCAPLUS
- (9) Mitsubishi Chemical Ind; JP 53021197 A 1978 HCAPLUS
- (10) Richard, G; WO 9804559 A 1998 HCAPLUS
- (11) Sanofi Sa; EP 0156734 A 1985 HCAPLUS
- (12) Schering Ag; DE 19617862 A 1997 HCAPLUS
- (13) Tarzia, G; FARMACO EDIZIONE SCIENTIFICA 1988, V43(2), P189 HCAPLUS

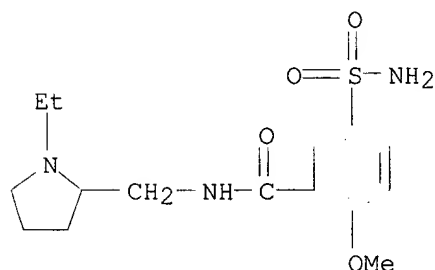
IT 15676-16-1, Sulpiride

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(triazolo-pyridazine deriv. GABAA ligands and therapeutic use, alone or with other compds.)

RN 15676-16-1 HCAPLUS

CN Benzamide, 5-(aminosulfonyl)-N-[(1-ethyl-2-pyrrolidinyl)methyl]-2-methoxy-(9CI) (CA INDEX NAME)



L127 ANSWER 20 OF 28 HCAPLUS COPYRIGHT 2003 ACS

AN 1999:64683 HCAPLUS

DN 130:125258

TI Highly selective **butyrylcholinesterase** inhibitors for the treatment and diagnosis of **Alzheimer's** disease and **dementias**

IN Greig, Nigel H.; Yu, Qian-sheng; Brossi, Arnold; Soncrant, Timothy T.; Hausman, Marvin

PA Axonyx, USA; National Institute of Health

SO PCT Int. Appl., 50 pp.  
CODEN: PIXXD2

DT **Patent**

LA English

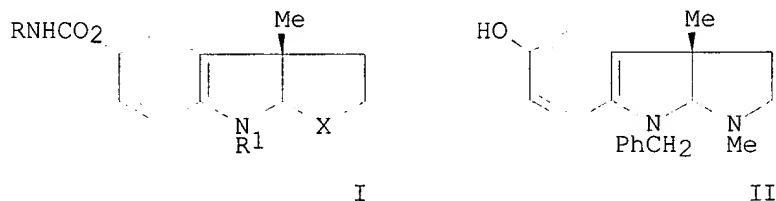
IC ICM A61K031-40  
ICS C07D209-58; C07D491-04; C07D491-048

CC 31-5 (Alkaloids)  
Section cross-reference(s): 1

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 9902154	A1	19990121	WO 1998-US14063	19980709 <--
	W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, GM, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
	RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
	AU 9882931	A1	19990208	AU 1998-82931	19980709 <--
	AU 749088	B2	20020620		
	EP 949920	A1	19991020	EP 1998-933230	19980709 <--
	R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO				
	JP 2001500165	T2	20010109	JP 1999-508849	19980709 <--
	MX 9902293	A	20000731	MX 1999-2293	19990309 <--
	US 6410747	B1	20020625	US 1999-254494	19990617 <--
	US 2002094999	A1	20020718	US 2002-71488	20020207 <--
PRAI	US 1997-52087P	P	19970709 <--		
	WO 1998-US14063	W	19980709 <--		
	US 1999-254494	A3	19990617 <--		

GI



AB The present disclosure relates to the discovery that highly selective butyrylcholinesterase inhibitors I [R = Me, Ph, C<sub>6</sub>H<sub>4</sub>Me-2, C<sub>6</sub>H<sub>4</sub>(CHMe<sub>2</sub>)-4, C<sub>6</sub>H<sub>4</sub>OMe-4, C<sub>6</sub>H<sub>4</sub>Me-4, C<sub>6</sub>H<sub>4</sub>(CNMe<sub>2</sub>)-2; R<sub>1</sub> = H, Me, CH<sub>2</sub>Ph; X = NH, NMe, NCH<sub>2</sub>Ph, O, S] can prevent or treat **cognitive** impairments assocd. with aging or **Alzheimer's** disease wherein said butyrylcholinesterase inhibitor has a selectivity ratio of butyrylcholinesterase inhibition to acetylcholinesterase inhibition of greater than about 15 to 1. A preferred butyrylcholinesterase inhibitor is N8-benzylnorcymserine [I; R = C<sub>6</sub>H<sub>4</sub>(CHMe<sub>2</sub>)-4, R<sub>1</sub> = CH<sub>2</sub>Ph, X = NMe] and is prepd. via reaction of hexahydropyrrolo[2,3-b]indol-5-ol II with 4-(Me<sub>2</sub>CH)C<sub>6</sub>H<sub>4</sub>NCO in Et<sub>2</sub>O.

Cymserine showed protection against a cholinergic forebrain lesion-induced increase in .beta.-amyloid precursor protein and improves cognitive performance in rats at 05.mg/kg.

- ST **Alzheimer** disease treatment cymserine deriv analog;  
**dementia** treatment cymserine deriv analog; butyrylcholinesterase inhibitor diagnosis **Alzheimer** disease; benzylnorcymserine prepn treatment **Alzheimer** disease; **amyloid** precursor protein secretion inhibitor cymserine
- IT **Mental disorder**  
 (dementia; selective butyrylcholinesterase inhibitors for the treatment and diagnosis of **Alzheimer's** disease and dementias)
- IT Aging, animal  
 (disorder, senility; selective butyrylcholinesterase inhibitors for the treatment and diagnosis of **Alzheimer's** disease and dementias)
- IT **Alzheimer's disease**  
 (selective butyrylcholinesterase inhibitors for the treatment and diagnosis of **Alzheimer's** disease and dementias)
- IT Aging, animal  
 (senility; selective butyrylcholinesterase inhibitors for the treatment and diagnosis of **Alzheimer's** disease and dementias)
- IT **Amyloid precursor proteins**  
 RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)  
 (.beta.-, secretion inhibitors; selective butyrylcholinesterase inhibitors for the treatment and diagnosis of **Alzheimer's** disease and dementias)
- IT 219920-75-9P, N8-Benzylnorcymserine 219920-88-4P, N1,N8-Bisbenzynorcymserine  
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)  
 (selective butyrylcholinesterase inhibitors for the treatment and diagnosis of **Alzheimer's** disease and dementias)
- IT 219920-78-2P, N8-Norcymserine 219920-81-7P, N1,N8-Bisnorcymserine  
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)  
 (selective butyrylcholinesterase inhibitors for the treatment and diagnosis of **Alzheimer's** disease and dementias)
- IT **57-47-6**, Physostigmine 6091-05-0, Physovenine 6091-57-2, Eseramine 19573-10-5, N8-Norphysostigmine 101246-66-6, Phenserine 114546-08-6, N1-Phenethylnorphysostigmine 116103-17-4, N1-Benzylnorphysostigmine 116103-18-5, N1-Norphysostigmine 116103-19-6, N1-Allylnorphysostigmine 116979-38-5, 4'-Methoxyphenserine 136092-41-6, Phensvenine 136092-42-7, Cymssvenine 145209-30-9, Tolserine 145209-31-0, N1-Benzylnortolserine 145209-32-1, N1-Nortolserine 145209-33-2, Tolsvenine 145209-34-3, 4'-Methylphenserine 145209-38-7, 2'-Isopropylphenserine 145209-39-8, Cymserine 145209-39-8D, carbon-11-labeled 145209-43-4, N1-Benzylnorphenserine 145209-44-5, N1-Norphenserine 145209-45-6, N1-Benzylnorcymserine 145209-46-7, N1-Norcymserine 145209-49-0, Thiaphenserine 145209-50-3, Thiatolserine 145209-51-4, Thiacymsenine 145237-06-5, Thiaphysovenine 171075-53-9, N1-Phenethylnorphenserine 171075-56-2, N1-Phenethylnortolserine 193604-43-2, N8-Benzylnorphysostigmine 193604-44-3, N8-Benzylnorphenserine 193604-45-4, N8-Norphenserine 207729-61-1, N1,N8-Bisnorphenserine 207729-62-2, N1,N8-Bisnorphysostigmine 207729-68-8, N1,N8-Bisbenzylnorphenserine 207729-69-9, N1,N8-Bisbenzylnorphysostigmine 219920-69-1, N1-Phenethylnorcymserine 219921-12-7, N8-Benzylnortolserine



219921-17-2, N8-Nortolserine 219921-23-0, N1,N8-Bisbenzylnortolserine

219922-89-1, N1,N8-Bisnortolserine

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); **THU (Therapeutic use)**; BIOL (Biological study); USES (Uses)

(selective butyrylcholinesterase inhibitors for the treatment and diagnosis of **Alzheimer's** disease and **dementias**)

IT 9000-81-1, Acetylcholinesterase 9001-08-5, Butyrylcholinesterase  
RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)

(selective butyrylcholinesterase inhibitors for the treatment and diagnosis of **Alzheimer's** disease and **dementias**)

IT 31027-31-3, 4-Isopropylphenyl isocyanate 193604-42-1,  
(-)-(3aS)-8-Benzyl-1,3a-dimethyl-1,2,3,3a,8,8a-hexahydropyrrolo[2,3-b]indol-5-ol 207729-66-6, (-)-(3aS)-1,8-Dibenzyl-3a-methyl-1,2,3,3a,8,8a-hexahydro-5-methoxypyrrolo[2,3-b]indole 207729-67-7,  
(-)-(3aS)-1,8-Dibenzyl-3a-methyl-1,2,3,3a,8,8a-hexahydropyrrolo[2,3-b]indol-5-ol

RL: RCT (Reactant); RACT (Reactant or reagent)

(selective butyrylcholinesterase inhibitors for the treatment and diagnosis of **Alzheimer's** disease and **dementias**)

RE.CNT 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD

RE

- (1) Brossi; US 5171750 A 1992 HCAPLUS
- (2) Brossi; US 5378723 A 1995 HCAPLUS
- (3) Greig; US 5409948 A 1995 HCAPLUS
- (4) Hamer; US 5541216 A 1996 HCAPLUS
- (5) Touvinen; Toxicol Appl Pharmacol 1996, V140(2), P364

IT 57-47-6, Physostigmine

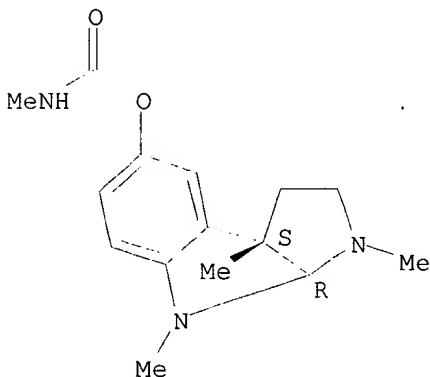
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); **THU (Therapeutic use)**; BIOL (Biological study); USES (Uses)

(selective butyrylcholinesterase inhibitors for the treatment and diagnosis of **Alzheimer's** disease and **dementias**)

RN 57-47-6 HCAPLUS

CN Pyrrolo[2,3-b]indol-5-ol, 1,2,3,3a,8,8a-hexahydro-1,3a,8-trimethyl-, methylcarbamate (ester), (3aS,8aR)- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).



L127 ANSWER 21 OF 28 HCAPLUS COPYRIGHT 2003 ACS

AN 1998:239104 HCAPLUS

DN 128:279000

TI Method for the treatment of **neurological** or **neuropsychiatric** disorders using therapy which blocks and/or inhibits melatonin, precursors thereof, and/or metabolic products thereof.

IN Willis, Gregory Lynn  
 PA Willis, Gregory Lynn, Australia  
 SO PCT Int. Appl., 55 pp.  
 CODEN: PIXXD2  
 DT **Patent**  
 LA English  
 IC ICM A61K031-135  
 ICS A61K031-165; A61N005-06; A61B017-00  
 CC 1-11 (Pharmacology)  
 Section cross-reference(s): 2

FAN.CNT 4

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 9815267	A1	19980416	WO 1997-AU661	19971003 <--
	W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
	RW: GH, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG				
	AU 9743725	A1	19980505	AU 1997-43725	19971003 <--
	AU 736005	B2	20010726		
	EP 964679	A1	19991222	EP 1997-941747	19971003 <--
	R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO				
	JP 2001503394	T2	20010313	JP 1998-517016	19971003 <--
	US 6310085	B1	20011030	US 1999-285859	19990402 <--
	US 2002068692	A1	20020606	US 2001-971783	20011009 <--
PRAI	AU 1996-2745	A	19961004	<--	
	WO 1997-AU661	W	19971003	<--	
	US 1999-285859	A2	19990402	<--	
AB	A method for the treatment and/or prophylaxis of a <b>neurol.</b> or <b>neuropsychiatric</b> disorder assocd. with altered dopamine function comprises subjecting a patient in need thereof to therapy which blocks and/or inhibits melatonin, precursors thereof and/or metabolic products thereof.				
ST	melatonin inhibition <b>neurol neuropsychiatric</b> disorder				
IT	<b>Brain, disease</b> (Gilles de la Tourette syndrome; <b>neurol.</b> or <b>neuropsychiatric</b> disorder treatment using therapy which blocks and/or inhibits melatonin, precursors thereof, and/or metabolic products thereof.)				
IT	<b>Nervous system</b> (Huntington's chorea; <b>neurol.</b> or <b>neuropsychiatric</b> disorder treatment using therapy which blocks and/or inhibits melatonin, precursors thereof, and/or metabolic products thereof.)				
IT	<b>Mental disorder</b> (Pick's disease; <b>neurol.</b> or <b>neuropsychiatric</b> disorder treatment using therapy which blocks and/or inhibits melatonin, precursors thereof, and/or metabolic products thereof.)				
IT	Stress, animal (acute stress disorder; <b>neurol.</b> or <b>neuropsychiatric</b> disorder treatment using therapy which blocks and/or inhibits melatonin, precursors thereof, and/or metabolic products thereof.)				
IT	<b>Mental disorder</b> (agoraphobia; <b>neurol.</b> or <b>neuropsychiatric</b> disorder treatment using therapy which blocks and/or inhibits melatonin, precursors thereof, and/or metabolic products thereof.)				
IT	<b>Nervous system</b> (akathisia; <b>neurol.</b> or <b>neuropsychiatric</b> disorder				

- treatment using therapy which blocks and/or inhibits melatonin, precursors thereof, and/or metabolic products thereof.)
- IT Appetite  
(anorexia **nervosa**; **neurol.** or **neuropsychiatric** disorder treatment using therapy which blocks and/or inhibits melatonin, precursors thereof, and/or metabolic products thereof.)
- IT Melatonin receptors  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(antagonists; **neurol.** or **neuropsychiatric** disorder treatment using therapy which blocks and/or inhibits melatonin, precursors thereof, and/or metabolic products thereof.)
- IT Anorexia  
(cachexia; **neurol.** or **neuropsychiatric** disorder treatment using therapy which blocks and/or inhibits melatonin, precursors thereof, and/or metabolic products thereof.)
- IT Ion channel blockers  
(calcium; **neurol.** or **neuropsychiatric** disorder treatment using therapy which blocks and/or inhibits melatonin, precursors thereof, and/or metabolic products thereof.)
- IT Mental disorder  
(**dementia**; **neurol.** or **neuropsychiatric** disorder treatment using therapy which blocks and/or inhibits melatonin, precursors thereof, and/or metabolic products thereof.)
- IT Mental disorder  
(depression, anxiety disorders due to; **neurol.** or **neuropsychiatric** disorder treatment using therapy which blocks and/or inhibits melatonin, precursors thereof, and/or metabolic products thereof.)
- IT Nervous system  
(dystonia, acute; **neurol.** or **neuropsychiatric** disorder treatment using therapy which blocks and/or inhibits melatonin, precursors thereof, and/or metabolic products thereof.)
- IT Antiparkinsonian agents  
(including for **neuroleptic-induced Parkinsonism**; **neurol.** or **neuropsychiatric** disorder treatment using therapy which blocks and/or inhibits melatonin, precursors thereof, and/or metabolic products thereof.)
- IT Behavior  
(locomotor; **neurol.** or **neuropsychiatric** disorder treatment using therapy which blocks and/or inhibits melatonin, precursors thereof, and/or metabolic products thereof.)
- IT Disease, animal  
(malignant syndrome; **neurol.** or **neuropsychiatric** disorder treatment using therapy which blocks and/or inhibits melatonin, precursors thereof, and/or metabolic products thereof.)
- IT Diagnosis  
(melatonin in **neurol.** or **neuropsychiatric** disorder diagnosis)
- IT Anti-Alzheimer's agents  
Anxiolytics  
Body weight  
Drug delivery systems  
Multiple sclerosis  
Nervous system agents  
Phototherapy  
Psychotropics  
Schizophrenia  
Wernicke-Korsakoff syndrome  
(**neurol.** or **neuropsychiatric** disorder treatment using therapy which blocks and/or inhibits melatonin, precursors thereof, and/or metabolic products thereof.)
- IT Mental disorder

- (obsession-compulsion; **neurol. or neuropsychiatric** disorder treatment using therapy which blocks and/or inhibits melatonin, precursors thereof, and/or metabolic products thereof.)
- IT Anxiety  
(panic disorder; **neurol. or neuropsychiatric** disorder treatment using therapy which blocks and/or inhibits melatonin, precursors thereof, and/or metabolic products thereof.)
- IT Movement disorders  
(periodic limb movement syndrome and others; **neurol. or neuropsychiatric** disorder treatment using therapy which blocks and/or inhibits melatonin, precursors thereof, and/or metabolic products thereof.)
- IT **Mental disorder**  
(post-traumatic stress disorder; **neurol. or neuropsychiatric** disorder treatment using therapy which blocks and/or inhibits melatonin, precursors thereof, and/or metabolic products thereof.)
- IT **Brain, disease**  
(**stroke**; **neurol. or neuropsychiatric** disorder treatment using therapy which blocks and/or inhibits melatonin, precursors thereof, and/or metabolic products thereof.)
- IT Paralysis  
(subnuclear, progressive; **neurol. or neuropsychiatric** disorder treatment using therapy which blocks and/or inhibits melatonin, precursors thereof, and/or metabolic products thereof.)
- IT Pineal gland  
(surgical ablation or destruction; **neurol. or neuropsychiatric** disorder treatment using therapy which blocks and/or inhibits melatonin, precursors thereof, and/or metabolic products thereof.)
- IT **Nervous system**  
(tardive dyskinesia; **neurol. or neuropsychiatric** disorder treatment using therapy which blocks and/or inhibits melatonin, precursors thereof, and/or metabolic products thereof.)
- IT Anti-ischemic agents  
(trans-ischemic attack; **neurol. or neuropsychiatric** disorder treatment using therapy which blocks and/or inhibits melatonin, precursors thereof, and/or metabolic products thereof.)
- IT Drugs  
(veterinary; **neurol. or neuropsychiatric** disorder treatment using therapy which blocks and/or inhibits melatonin, precursors thereof, and/or metabolic products thereof.)
- IT Adrenoceptor antagonists  
(.beta.-; **neurol. or neuropsychiatric** disorder treatment using therapy which blocks and/or inhibits melatonin, precursors thereof, and/or metabolic products thereof.)
- IT 51-61-6, Dopamine, biological studies  
RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)  
(altered dopamine function; **neurol. or neuropsychiatric** disorder treatment using therapy which blocks and/or inhibits melatonin, precursors thereof, and/or metabolic products thereof.)
- IT 1199-18-4 28289-54-5, MPTP  
RL: ADV (Adverse effect, including toxicity); BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study)  
(**neurol. or neuropsychiatric** disorder treatment using therapy which blocks and/or inhibits melatonin, precursors thereof, and/or metabolic products thereof.)
- IT 9002-79-3, Melanocyte-stimulating hormone 29122-68-7, Atenolol  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (**Therapeutic use**); BIOL (Biological

study); USES (Uses)

(**neurol.** or **neuropsychiatric** disorder treatment using therapy which blocks and/or inhibits melatonin, precursors thereof, and/or metabolic products thereof.)

IT 73-31-4, Melatonin 73-31-4D, Melatonin, precursors and metabolic products

RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)

(**neurol.** or **neuropsychiatric** disorder treatment using therapy which blocks and/or inhibits melatonin, precursors thereof, and/or metabolic products thereof.)

RE.CNT 15 THERE ARE 15 CITED REFERENCES AVAILABLE FOR THIS RECORD

RE

- (1) Adler, L; Psychopharmacology Bulletin 1991, V27(2), P107 MEDLINE
- (2) Anon; EP 146113 HCAPLUS
- (3) Anon; The Merck Manual of Diagnosis and Therapy, 16th Edition 1992, P1499
- (4) Artemenko, A; 1996
- (5) Chuprikov; US 5137018 1992
- (6) Dubocovich; US 5093352 1992 HCAPLUS
- (7) Dubocovich; US 5283343 1994 HCAPLUS
- (8) Garcia-Talavera, B; 1984
- (9) Glaxo Group Limited; WO 9529173 1995 HCAPLUS
- (10) Koller, W; Arch Neurol 1987, V44, P921 MEDLINE
- (11) Martindale; The Extra Pharmacopoeia 28th Edition 1982, P1337
- (12) Miles, A; Biol Psychiatry 1988, V23, P405 HCAPLUS
- (13) Searfoss; US 5046494 1991
- (14) Sherer, M; Neurosci Lett 1985, V58(3), P277 MEDLINE
- (15) Wilbur, R; Prog Neuro-Psychopharmacol and Biol Psychiat 1988, V12, P848

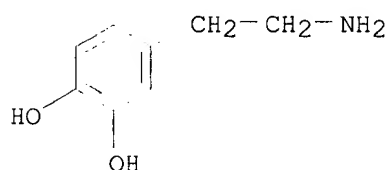
IT 51-61-6, Dopamine, biological studies

RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)

(altered dopamine function; **neurol.** or **neuropsychiatric** disorder treatment using therapy which blocks and/or inhibits melatonin, precursors thereof, and/or metabolic products thereof.)

RN 51-61-6 HCAPLUS

CN 1,2-Benzenediol, 4-(2-aminoethyl)- (9CI) (CA INDEX NAME)



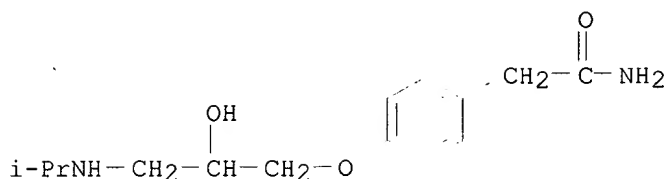
IT 29122-68-7, Atenolol

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); **THU (Therapeutic use)**; BIOL (Biological study); USES (Uses)

(**neurol.** or **neuropsychiatric** disorder treatment using therapy which blocks and/or inhibits melatonin, precursors thereof, and/or metabolic products thereof.)

RN. 29122-68-7 HCAPLUS

CN Benzeneacetamide, 4-[2-hydroxy-3-[(1-methylethyl)amino]propoxy]- (9CI) (CA INDEX NAME)



L127 ANSWER 22 OF 28 HCAPLUS COPYRIGHT 2003 ACS

AN 1998:161121 HCAPLUS

DN 128:213404

TI Sulfonyl fluorides for the treatment of **Alzheimer's** disease

IN Moss, Donald Eugene

PA Board of Regents of the University of Texas System, USA

SO PCT Int. Appl., 54 pp.

CODEN: PIXXD2

DT **Patent**

LA English

IC ICM A61K031-135

ICS A61K031-10

CC 1-11 (Pharmacology)

Section cross-reference(s): 63

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 9808497	A1	19980305	WO 1997-US15024	19970826 <--
	W: AU, CA, JP, MX				
	RW: AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				
	US 5798392	A	19980825	US 1996-705858	19960828 <--
	US 5798392	C1	20020604		
	AU 9740915	A1	19980319	AU 1997-40915	19970826 <--
	AU 731755	B2	20010405		
	EP 921790	A1	19990616	EP 1997-938628	19970826 <--
	R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI				
	JP 2000517317	T2	20001226	JP 1998-511856	19970826 <--
PRAI	US 1996-705858	A	19960828	<--	
	WO 1997-US15024	W	19970826	<--	
AB	A pharmaceutical compn. is provided which comprises a sulfonyl fluoride and a pharmaceutically acceptable carrier. Also provided is a method of treating <b>Alzheimer's</b> disease in an individual in need of such treatment, comprising administering to the individual a therapeutically ED of methanesulfonyl fluoride. Further provided is a method of enhancing <b>cognitive</b> performance in an individual in need of such treatment, comprising administering to the individual a therapeutically ED of methanesulfonyl fluoride.				
ST	sulfonyl fluoride <b>Alzheimer</b> disease; methanesulfonyl fluoride <b>Alzheimer</b> disease; <b>cognition</b> enhancer methanesulfonyl fluoride				
IT	<b>Parkinson's disease</b> <b>Parkinson's disease</b> (Guamanian <b>parkinsonism-dementia</b> ; sulfonyl fluorides for <b>Alzheimer's</b> disease treatment)				
IT	Erythrocyte (acetylcholinesterase; sulfonyl fluorides for <b>Alzheimer's</b> disease treatment)				
IT	<b>Nervous system</b> (central, disease, acetylcholine insufficiency-related; sulfonyl fluorides for <b>Alzheimer's</b> disease treatment)				
IT	<b>Mental disorder</b>				

(**dementia**, Boxer's; sulfonyl fluorides for **Alzheimer**'s disease treatment)

IT **Anti-Alzheimer's agents**  
**Antiparkinsonian agents**  
**Cognition enhancers**  
Drug delivery systems  
Enzyme kinetics  
Michaelis constant  
(sulfonyl fluorides for **Alzheimer's** disease treatment)

IT **Lecithins**  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(sulfonyl fluorides with other agents for **Alzheimer's** disease treatment)

IT **Sulfonyl halides**  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(sulfonyl fluorides; sulfonyl fluorides for **Alzheimer's** disease treatment)

IT 328-86-9 329-98-6, Phenylmethanesulfonyl fluoride 368-43-4, Benzenesulfonyl fluoride 368-72-9, 3-Amino-4-chlorobenzenesulfonyl fluoride 455-16-3, p-Toluenesulfonyl fluoride 498-74-8 558-25-8, Methanesulfonyl fluoride 754-03-0, Ethanesulfonyl fluoride 63805-73-2, 2-Propanesulfonyl fluoride 204260-09-3  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(sulfonyl fluorides for **Alzheimer's** disease treatment)

IT 51-84-3, Acetylcholine, biological studies 9000-81-1, Acetylcholinesterase 9001-08-5, Butyrylcholinesterase  
RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)  
(sulfonyl fluorides for **Alzheimer's** disease treatment)

IT **504-24-5**, 4-Aminopyridine 3576-73-6, RS86  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); **THU (Therapeutic use)**; BIOL (Biological study); USES (Uses)  
(sulfonyl fluorides with other agents for **Alzheimer's** disease treatment)

RE.CNT 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD  
RE  
(1) Moss, D; Adv Behav Biol 1986, V29, P551 HCAPLUS  
(2) Moss, D; Curr Res Alzheimer Ther:Cholinesterase Inhib 1988, P305 HCAPLUS  
(3) Palacios-Esquivel, R; Neurobiol Aging 1993, V14(1), P93 HCAPLUS

IT **504-24-5**, 4-Aminopyridine  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); **THU (Therapeutic use)**; BIOL (Biological study); USES (Uses)  
(sulfonyl fluorides with other agents for **Alzheimer's** disease treatment)

RN 504-24-5 HCAPLUS  
CN 4-Pyridinamine (9CI) (CA INDEX NAME)



L127 ANSWER 23 OF 28 HCAPLUS COPYRIGHT 2003 ACS

AN 1998:147210 HCAPLUS

DN 128:201063

TI Cholinesterase inhibitors for treatment of **Parkinson's** disease

IN Hutchinson, Michael

PA New York University, USA

SO PCT Int. Appl., 26 pp.

CODEN: PIXXD2

DT **Patent**

LA English

IC ICM A61K031-55

ICS A61K031-40; A61K031-44; A61K031-505; A61K031-66; A61K031-195

CC 1-11 (Pharmacology)

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 9807431	A1	19980226	WO 1997-US14684	19970821 <--
	W: AU, CA, IL, JP, MX				
	RW: AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				
	AU 9740789	A1	19980306	AU 1997-40789	19970821 <--
	US 5965571	A	19991012	US 1997-915736	19970821 <--
PRAI	US 1996-22746P	P	19960822 <--		
	WO 1997-US14684	W	19970821 <--		
AB	<b>Parkinson's</b> disease can be treated with at least one cholinesterase inhibitor. The cholinesterase inhibitor has been found to alleviate both any symptoms of <b>dementia</b> as well as to reduce rigidity and improve motor function. E.g., on administration of 40 mg tacrine, building up to 60 mg/day after seven weeks, a patient exhibited remarkable improvements in <b>dementia</b> and rigidity.				
ST	cholinesterase inhibitor <b>Parkinsons</b> disease				
IT	<b>Parkinson's</b> disease (cholinesterase inhibitors for treatment of <b>Parkinson's</b> disease)				
IT	<b>Mental disorder</b> ( <b>dementia</b> ; cholinesterase inhibitors for treatment of <b>Parkinson's</b> disease)				
IT	321-64-2, Tacrine RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (cholinesterase inhibitors for treatment of <b>Parkinson's</b> disease)				
IT	59-92-7, Levodopa, biological studies RL: MOA (Modifier or additive use); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (cholinesterase inhibitors for treatment of <b>Parkinson's</b> disease)				
IT	52-68-6, Metrifonate 57-47-6, Physostigmine 357-70-0, Galanthamine 987-78-0, Citicoline 101246-68-8, Heptastigmine 118909-22-1, Velnacrine maleate RL: <b>THU (Therapeutic use)</b> ; BIOL (Biological study); USES (Uses) (cholinesterase inhibitors for treatment of <b>Parkinson's</b> disease)				
IT	9000-81-1, Acetylcholinesterase 9001-08-5, Cholinesterase RL: BSU (Biological study, unclassified); BIOL (Biological study) (inhibitors; cholinesterase inhibitors for treatment of <b>Parkinson's</b> disease)				

RE.CNT 2 THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD

RE

(1) Boar; US 5585378 A 1996 HCAPLUS

(2) Rosin; US 4948807 A 1990 HCAPLUS



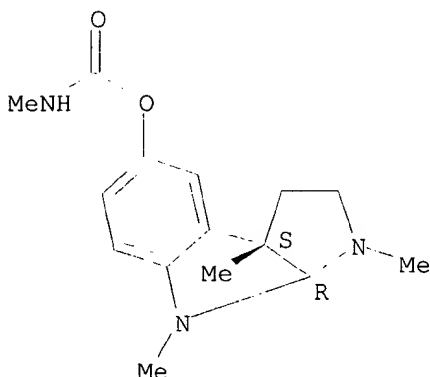
IT 57-47-6, Physostigmine

RL: **THU (Therapeutic use)**; BIOL (Biological study); USES (Uses)  
(cholinesterase inhibitors for treatment of **Parkinson's**  
disease)

RN 57-47-6 HCAPLUS

CN Pyrrolo[2,3-b]indol-5-ol, 1,2,3,3a,8,8a-hexahydro-1,3a,8-trimethyl-,  
methylcarbamate (ester), (3aS,8aR)- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).



L127 ANSWER 24 OF 28 HCAPLUS COPYRIGHT 2003 ACS

AN 1997:617007 HCAPLUS

DN 127:288186

TI Methods of treating **neurological** diseases and etiologically  
related symptomology using carbonyl trapping agents in combination with  
previously known medicaments

IN Shapiro, Howard K.

PA USA

SO U.S., 37 pp., Cont.-in-part of U.S. Ser. No. 26,617, abandoned.

CODEN: USXXAM

DT **Patent**

LA English

IC ICM A01N043-04

ICS A01N061-00; C07H001-00; C08B037-08

NCL 514055000

CC 1-11 (Pharmacology)

Section cross-reference(s): 63

FAN.CNT 3

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	US 5668117	A	19970916	US 1993-62201	19930629 <--
	CA 2166383	AA	19950112	CA 1994-2166383	19940628 <--
	WO 9501096	A1	19950112	WO 1994-US7277	19940628 <--
	W: AU, CA, JP				
	RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				
	AU 9472144	A1	19950124	AU 1994-72144	19940628 <--
	AU 692454	B2	19980611		
	EP 707446	A1	19960424	EP 1994-921405	19940628 <--
	R: DE, FR, GB, IT				
	JP 08512055	T2	19961217	JP 1994-503597	19940628 <--
PRAI	US 1991-660561	B1	19910222	<--	
	US 1993-26617	B2	19930223	<--	
	US 1993-62201	A	19930629	<--	
	WO 1994-US7277	W	19940628	<--	
OS	MARPAT 127:288186				
AB	Therapeutic compns. comprising an effective amt. of at least one carbonyl				

trapping agent alone or in combination with a therapeutically effective of a co-agent or medicament are disclosed. The compns. are used to treat a mammal suffering from a **neurol.** disease characterized by covalent bond crosslinking between the **nerve** cells, other cellular structures and their intracellular and extracellular components, with disease-induced carbonyl-contg. aliph. or arom. hydrocarbons present in mammals.

ST carbonyl trap drug combination **neurol** disease

IT **Nervous system**

(Huntington's chorea; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)

IT Glutamate antagonists

(NMDA antagonists; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)

IT Sulfhydryl group

(agents contg.; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)

IT **Nervous system**

(amyotrophic lateral sclerosis; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)

IT Antiarteriosclerotics

(antiatherosclerotics; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)

IT Sequestering agents

(bile acid; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)

IT Amines, biological studies

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(biogenic; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)

IT Ion channel blockers

(calcium; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)

IT 5-HT antagonists

Aging, animal

**Alzheimer's disease**

Analgesics

Anti-ischemic agents

Antiarrhythmics

Anticonvulsants

Antidepressants

**Antidiabetic agents**

Antihistamines

Antihypertensives

Antioxidants

Anxiolytics

Carbonyl group

Cholinergic agonists

Cholinergic antagonists

**Cognition enhancers**

Dopamine agonists

Drug delivery systems

Drug interactions

Hypolipemic agents

Immunosuppressants

Multiple sclerosis

**Nervous system agents**

**Parkinson's disease**

Platelet aggregation inhibitors

Psychotropics

Radical scavengers

Tranquilizers

Vasodilators

(carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)

IT Carbonyl compounds (organic), biological studies

RL: ADV (Adverse effect, including toxicity); BIOL (Biological study)

(carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)

IT Corticosteroids, biological studies

Hormones, animal, biological studies

Interferons

Lecithins

Phosphatidylcholines, biological studies

Vitamins

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)IT **Nerve**(conduction; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)IT **Drugs**

Kidney

(conjugating agents facilitating kidney drug elimination; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)IT **Animal cell**(crosslinking; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)IT **Gastric emptying**(delayed; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)IT **Nerve, disease**(demyelination, urinary incontinence in; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)IT **Ear**(disease, tinnitus; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)IT **Nervous system**(disease; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)IT **Neurotransmission**(enhancers; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)IT **Proteins, general, biological studies**

RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)

(fibroblast; **neurofilament** crosslinking in chromosome 17 hereditary sensory and motor **neuropathy**)IT **Digestive tract**(gastroesophageal reflux; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)IT **Chromosome**(human 17, hereditary sensory and motor **neuropathy**; **neurofilament** crosslinking in chromosome 17 hereditary sensory

- and motor **neuropathy**)
- IT Bladder
  - (incontinence, from **Alzheimer's senile dementia** or other disease; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)
- IT Heart, disease
  - (ischemia, agents for treatment of; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)
- IT Brain
  - (metab., enhancers; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)
- IT Metabolism
  - (metabolites at risk of depletion; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)
- IT Gangliosides
  - RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
  - (mixed cow brain; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)
- IT Kidney, disease
  - (nephrotic syndrome, diabetes-related, agents for treatment of; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)
- IT Crosslinking
  - (**nerve** cell; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)
- IT Fibroblast
  - (**neurofilament** crosslinking in chromosome 17 hereditary sensory and motor **neuropathy**)
- IT Cytoskeleton
  - (**neurofilament**; **neurofilament** crosslinking in chromosome 17 hereditary sensory and motor **neuropathy**)
- IT Nerve
  - (**neuron**, crosslinking; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)
- IT Nerve, disease
  - (**neuropathy**, hereditary motor and sensory; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)
- IT Anti-inflammatory agents
  - (nonsteroidal; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)
- IT Nerve, disease
  - (**peripheral neuropathy**, urinary incontinence in; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)
- IT Carbohydrates, biological studies
  - RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
  - (plant, non-digestible, edible; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)
- IT Polymers, biological studies
  - RL: BAC (Biological activity or effector, except adverse); BSU (Biological

- study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(polyamine-related; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)
- IT Amines, biological studies  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(polymers; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)
- IT **Nerve, disease**  
(**polyneuropathy**, alc.; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)
- IT Receptors  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(postsynaptic, agonists; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)
- IT Bile acids  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(sequestrants; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)
- IT **Nervous system**  
(spinocerebellar ataxia; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)
- IT Reagents  
RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(suspending; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)
- IT Polycyclic compounds  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(tricyclic; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)
- IT **Diabetes mellitus**  
(urinary incontinence in; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)
- IT Blood  
(viscosity, agents decreasing; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)
- IT Adrenoceptors  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(.alpha.-, .alpha.-adrenergic agents; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)
- IT Adrenoceptor antagonists  
(.beta.-; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)
- IT 9001-66-5  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(A and B, inhibitors; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)
- IT 51-84-3, Acetylcholine, biological studies  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(agents enhancing synthesis, storage, or release of; carbonyl trapping agent combination with other drug for treatment of **neurol.**

- diseases and etiol. related symptomol.)
- IT 70-18-8, Glutathione, biological studies  
 RL: BSU (Biological study, unclassified); BIOL (Biological study)  
 (agents facilitating activity of; carbonyl trapping agent combination  
 with other drug for treatment of **neurol.** diseases and etiol.  
 related symptomol.)
- IT 50-18-0, Cyclophosphamide 54-96-6, 3,4-Diaminopyridine 56-40-6,  
 Glycine, biological studies 57-47-6, Physostigmine 58-56-0,  
 Pyridoxine hydrochloride 58-85-5, Biotin 59-02-9 59-30-3, Folic  
 acid, biological studies 59-43-8, Thiamine, biological studies  
 59-51-8, D,L-Methionine 59-67-6, Nicotinic acid, biological studies  
 59-92-7, Levodopa, biological studies 68-19-9, Vitamin B12 68-41-7,  
 D-Cycloserine 72-19-5, L-Threonine, biological studies 79-83-4,  
 Pantothenic acid 83-88-5, Vitamin B2, biological studies 98-92-0,  
 Nicotinamide 137-08-6 137-58-6, Lidocaine 150-13-0, p-Aminobenzoic  
 acid 302-79-4D, Retinoic acid, derivs. 302-84-1, Serine 321-64-2,  
 Tacrine 357-70-0, Galanthamine 364-62-5, Metoclopramide 446-86-6,  
 Azathioprine 456-59-7, Cyclandelate 504-24-5, 4-Aminopyridine  
 645-88-5, Aminoxyacetic acid 657-24-9, Metformin 768-94-5, Amantadine  
 1134-47-0, Baclofen 1195-16-0 1406-18-4, Vitamin E 3200-06-4,  
 Praxilene 3286-46-2, Sulbutiamine 4759-48-2, 13-cis-Retinoic acid  
 7235-40-7, .beta.-Carotene 7491-74-9, Piracetam 7782-49-2, Selenium,  
 biological studies 8059-24-3, Vitamin B6 9004-10-8D, Insulin, derivs.,  
 biological studies 11000-17-2D, Vasopressin, analogs 13345-51-2D,  
 Prostaglandin B1, oligomers 14611-51-9, Selegiline 15301-69-6,  
 Flavoxate 18601-90-6, Thiamine mononitrate 23210-56-2, Ifenprodil  
 24305-27-9, Thyrotropin releasing factor 28704-27-0 28860-95-9,  
 Carbidopa 37758-47-7, Ganglioside GM1 41708-72-9, Tocainide  
 51012-32-9, Tiapride 51037-30-0, Acipimox 51481-61-9,  
 Cimetidine 54143-55-4, Flecainide 59865-13-3, Cyclosporine  
 66357-35-5, Ranitidine 72432-10-1, Aniracetam 73590-58-6, Omeprazole  
 76824-35-6, Famotidine 81098-60-4, Cisapride 103878-84-8, Lazabemide  
 105431-72-9, Linopirdine 196966-12-8, Anfaccine  
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological  
 study, unclassified); THU (Therapeutic use); BIOL (Biological  
 study); USES (Uses)  
 (carbonyl trapping agent combination with other drug for treatment of  
**neurol.** diseases and etiol. related symptomol.)
- IT 63-74-1D, Sulfanilamide, derivs.  
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological  
 study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES  
 (Uses)  
 (hypoglycemic; carbonyl trapping agent combination with other drug for  
 treatment of **neurol.** diseases and etiol. related symptomol.)
- IT 9000-81-1, Acetylcholinesterase  
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological  
 study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES  
 (Uses)  
 (inhibitors; carbonyl trapping agent combination with other drug for  
 treatment of **neurol.** diseases and etiol. related symptomol.)
- IT 9001-08-5, Cholinesterase 9015-82-1 9027-22-9, Decarboxylase  
 9028-31-3, Aldose reductase 9028-35-7, 3-Hydroxy-3-methylglutaryl CoA  
 reductase  
 RL: BSU (Biological study, unclassified); BIOL (Biological study)  
 (inhibitors; carbonyl trapping agent combination with other drug for  
 treatment of **neurol.** diseases and etiol. related symptomol.)
- IT 50-67-9, Serotonin, biological studies  
 RL: BSU (Biological study, unclassified); BIOL (Biological study)  
 (reuptake inhibitors, antidepressants; carbonyl trapping agent  
 combination with other drug for treatment of **neurol.** diseases  
 and etiol. related symptomol.)
- IT 12794-10-4D, Benzodiazepine, derivs.  
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological

study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(tranquilizers; carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)

IT 57-47-6, Physostigmine 504-24-5, 4-Aminopyridine

51481-61-9, Cimetidine

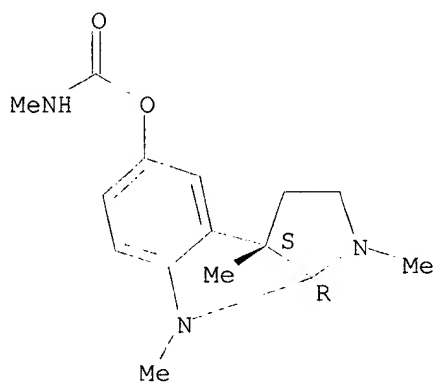
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); **THU (Therapeutic use)**; BIOL (Biological study); USES (Uses)

(carbonyl trapping agent combination with other drug for treatment of **neurol.** diseases and etiol. related symptomol.)

RN 57-47-6 HCAPLUS

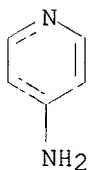
CN Pyrrolo[2,3-b]indol-5-ol, 1,2,3,3a,8,8a-hexahydro-1,3a,8-trimethyl-, methylcarbamate (ester), (3aS,8aR)- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).



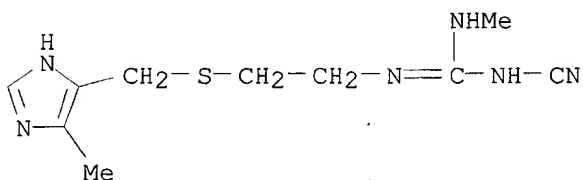
RN 504-24-5 HCAPLUS

CN 4-Pyridinamine (9CI) (CA INDEX NAME)



RN 51481-61-9 HCAPLUS

CN Guanidine, N-cyano-N'-methyl-N''-[2-[[[5-methyl-1H-imidazol-4-yl)methyl]thio]ethyl]- (9CI) (CA INDEX NAME)



L127 ANSWER 25 OF 28 HCAPLUS COPYRIGHT 2003 ACS

AN 1995:991057 HCAPLUS

DN 124:45735

TI Tacrine and cytochrome P450 oxidase inhibitors and methods of use for treatment of **Alzheimer's** disease

IN Woolf, Thomas F.

PA Warner Lambert Co., USA

SO U.S., 15 pp. Cont.-in-part of U.S. 5,422,350.

CODEN: USXXAM

DT **Patent**

LA English

IC ICM A61K031-44

ICS A61K031-415; A61K031-505; A61K031-495

NCL 514297000

CC 1-11 (Pharmacology)

FAN.CNT 2

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	US 5466696	A	19951114	US 1993-100917	19930809 <--
	US 5422350	A	19950606	US 1992-943323	19920910 <--
	WO 9405296	A2	19940317	WO 1993-US8459	19930908 <--
	WO 9405296	A3	19940428		
	W: AU, CA, CZ, FI, HU, JP, KR, NO, NZ, PT, RU, SK, UA				
	RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				
	EP 659082	A1	19950628	EP 1993-923110	19930908 <--
	EP 659082	B1	20020206		
	R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LI, LU, MC, NL, PT, SE				
	HU 70979	A2	19951128	HU 1995-704	19930908 <--
	HU 217845	B	20000428		
	JP 08501105	T2	19960206	JP 1994-507526	19930908 <--
	AU 669586	B2	19960613	AU 1993-52905	19930908 <--
	AU 9352905	A1	19940329		
	NZ 280680	A	20010831	NZ 1993-280680	19930908 <--
	AT 212845	E	20020215	AT 1993-923110	19930908 <--
	ES 2171419	T3	20020916	ES 1993-923110	19930908 <--
	FI 9501024	A	19950306	FI 1995-1024	19950306 <--
	NO 9500909	A	19950309	NO 1995-909	19950309 <--
PRAI	US 1992-943323	A2	19920910	<--	
	US 1993-100917	A	19930809	<--	
	NZ 1993-256940	A1	19930908	<--	
	WO 1993-US8459	W	19930908	<--	

OS MARPAT 124:45735

AB A method for treating **Alzheimer's** disease in a patient comprises administering to the patient an effective amt. of tacrine in combination with a P 450 1A2 oxidase inhibitor. Preferably, the inhibitor is a heterocyclic guanidine. The in vivo and in vitro metab. of tacrine and a proposed pathway for irreversible binding of tacrine to human liver microsomal protein are also included. Enoxacin, a specific P 450 1A2 inhibitor, not only decreased the rate of irreversible binding but also inhibited the overall rate of tacrine biotransformation.

ST cytochrome P450 oxidase inhibitor tacrine **Alzheimer**; P450 oxidase inhibitor tacrine **Alzheimer** disease

IT **Mental disorder**

(**Alzheimer's** disease, tacrine and cytochrome P 450 oxidase inhibitors and methods of use for treatment of **Alzheimer's** disease)

IT 9038-14-6

RL: BSU (Biological study, unclassified); BIOL (Biological study) (cytochrome P 450 1A2-dependent, inhibitors; tacrine and cytochrome P 450 oxidase inhibitors and methods of use for treatment of **Alzheimer's** disease)

IT 54-05-7, Chloroquine 64-99-3, Ethimizol 74011-58-8, Enoxacin 80288-49-9, Furafylline

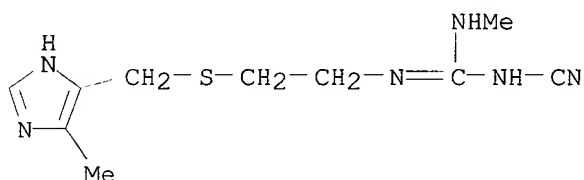
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study)

(tacrine and cytochrome P 450 oxidase inhibitors and methods of use for



*Harold  
Hendrick*

- treatment of **Alzheimer's** disease)
- IT 124027-47-0, 1-Hydroxytacrine 130073-98-2, 2-Hydroxytacrine  
130073-99-3, 4-Hydroxytacrine 136051-80-4  
RL: BOC (Biological occurrence); BSU (Biological study, unclassified); MFM (Metabolic formation); BIOL (Biological study); FORM (Formation, nonpreparative); OCCU (Occurrence)  
(tacrine and cytochrome P 450 oxidase inhibitors and methods of use for treatment of **Alzheimer's** disease)
- IT 321-64-2, Tacrine  
RL: BPR (Biological process); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)  
(tacrine and cytochrome P 450 oxidase inhibitors and methods of use for treatment of **Alzheimer's** disease)
- IT 56-49-5, 3-Methylcholanthrene 120-58-1, Isosafrole 1746-01-6, 2,3,7,8-Tetrachlorodibenzo-p-dioxin 6051-87-2, .beta.-Naphthoflavone 9035-51-2, Cytochrome P 450, biological studies 11097-69-1, Aroclor 1254  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(tacrine and cytochrome P 450 oxidase inhibitors and methods of use for treatment of **Alzheimer's** disease)
- IT 51481-61-9 52378-41-3 52378-49-1 52378-50-4 52378-59-3  
RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(tacrine and cytochrome P 450 oxidase inhibitors and methods of use for treatment of **Alzheimer's** disease)
- IT 51481-61-9  
RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(tacrine and cytochrome P 450 oxidase inhibitors and methods of use for treatment of **Alzheimer's** disease)
- RN 51481-61-9 HCAPLUS
- CN Guanidine, N-cyano-N'-methyl-N''-[2-[[[(5-methyl-1H-imidazol-4-yl)methyl]thio]ethyl]- (9CI) (CA INDEX NAME)



- L127 ANSWER 26 OF 28 HCAPLUS COPYRIGHT 2003 ACS
- AN 1995:940662 HCAPLUS
- DN 124:45496
- TI Clinical pharmacokinetics of physostigmine in patients with **Alzheimer's** disease
- AU Asthana, Sanjay; Greig, Nigel H.; Hegedus, Lajos; Holloway, Harold H.; Raffaele, Kathleen C.; Schapiro, Mark B.; Soncrant, Timothy T.
- CS National Institute Aging, National Institutes Health, Bethesda, MD, USA
- SO Clinical Pharmacology and Therapeutics (St. Louis) (1995), 58(3), 299-309  
CODEN: CLPTAT; ISSN: 0009-9236
- PB Mosby-Year Book
- DT Journal
- LA English
- CC 1-11 (Pharmacology)
- AB Our objective to study the pharmacokinetic and pharmacodynamic properties of physostigmine in subjects with **Alzheimer's** disease. Plasma physostigmine concn. and butyrylcholinesterase inhibition were measured in blood samples collected during and after a single high-dose (1 to 1.5 mg for 45 to 60 min) and a sustained low-dose steady-state i.v. infusion in nine subjects with **Alzheimer's** disease. Escalating doses (0.5

to 25 mg/day) were administered during a 2-wk period. A dose (2 to 12 mg/day) that optimized **cognition** in each subject was identified and then administered in a randomized, double-blind, placebo-controlled crossover design for 1 wk. The elimination half-life of physostigmine was 16.4  $\pm$  3.2 (SE) minutes. Clearance and vol. of distribution were 7.7  $\pm$  0.9 (SE) L/min and 2.4  $\pm$  0.6 (SE) L/kg, resp. Butyrylcholinesterase inhibition half-life was 83.7  $\pm$  5.2 (SE) minutes. During sustained steady-state infusion, plasma physostigmine concn. ( $r$  = 0.95) and butyrylcholinesterase inhibition ( $r$  = 0.99) were linearly correlated with the dose. In five **cognitive** responders, the memory enhancement was significantly correlated ( $r$  = 0.86;  $p$  < 0.05) with butyrylcholinesterase inhibition. These results showed that, in **cognitive** responders, memory enhancement by physostigmine in **Alzheimer's** disease is correlated directly to the magnitude of plasma cholinesterase inhibition. Furthermore, during single-dose conditions, the dynamic half-life is five-fold longer than the kinetic half-life.

ST physostigmine **Alzheimer** disease

IT **Mental disorder**

(**Alzheimer's** disease, clin. pharmacokinetics of physostigmine in humans with **Alzheimer's** disease)

IT 57-47-6, Physostigmine

RL: ADV (Adverse effect, including toxicity); BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); **THU** (**Therapeutic use**); BIOL (Biological study); USES (Uses)  
(clin. pharmacokinetics of physostigmine in humans with **Alzheimer's** disease)

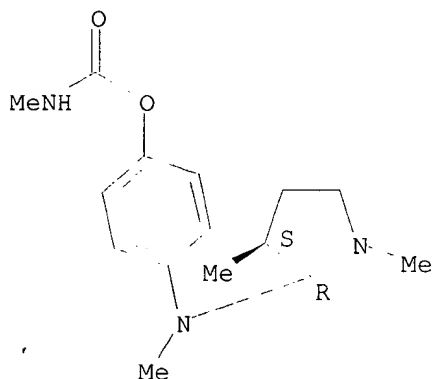
IT 57-47-6, Physostigmine

RL: ADV (Adverse effect, including toxicity); BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); **THU** (**Therapeutic use**); BIOL (Biological study); USES (Uses)  
(clin. pharmacokinetics of physostigmine in humans with **Alzheimer's** disease)

RN 57-47-6 HCAPLUS

CN Pyrrolo[2,3-b]indol-5-ol, 1,2,3,3a,8,8a-hexahydro-1,3a,8-trimethyl-, methylcarbamate (ester), (3aS,8aR)- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).



L127 ANSWER 27 OF 28 HCAPLUS COPYRIGHT 2003 ACS

AN 1995:723143 HCAPLUS

DN 123:102794

TI Pharmaceutical compositions and use thereof for treatment of **neurological** diseases and etiologically related symptomatology.

IN Shapiro, Howard K.

PA USA

SO PCT Int. Appl., 155 pp.

CODEN: PIXXD2

DT **Patent**

LA English

IC ICM A01N043-04

ICS A01N061-00; A61K031-73; C12Q001-68; G01N033-00; G01N033-539

CC 1-11 (Pharmacology)

FAN.CNT 3

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 9501096	A1	19950112	WO 1994-US7277	19940628 <--
	W: AU, CA, JP				
	RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				
	US 5668117	A	19970916	US 1993-62201	19930629 <--
	AU 9472144	A1	19950124	AU 1994-72144	19940628 <--
	AU 692454	B2	19980611		
	EP 707446	A1	19960424	EP 1994-921405	19940628 <--
	R: DE, FR, GB, IT				
	JP 08512055	T2	19961217	JP 1994-503597	19940628 <--
PRAI	US 1993-62201	A	19930629 <--		
	US 1991-660561	B1	19910222 <--		
	US 1993-26617	B2	19930223 <--		
	WO 1994-US7277	W	19940628 <--		
AB	<p>Pharmaceutical compns. for treatment of several <b>neurol.</b> diseases and pathophysiol.-related symptomol. in other body tissues, including peripheral <b>neuropathies</b>, secondary symptomol. of diabetes, <b>Alzheimer's</b> disease, <b>Parkinson's</b> disease, alc. <b>polyneuropathy</b> and age-onset symptomol., as well as analogous veterinary diseases, are disclosed. Spurious pathol. chem. crosslinking of normal intracellular structures is a fundamental aspect of these <b>neurol.</b> diseases. Covalent bond crosslinking of protein and lipid subcellular elements appear to underlie the formation of polyimd. <b>aggregates of neurofilaments</b> and other structural proteins, and lipofuscin. Pharmacol. intervention in some <b>neurol.</b> diseases using water-sol., small mol. wt. primary amines or their derivs. as oral therapeutic agents, may compete with cellular protein and lipid amine groups for reaction with disease-induced carbonyl-contg. aliph. and arom. hydrocarbons. Primary pharmacol. agents include 4-aminobenzoic acid and derivs. thereof to facilitate kidney recognition and removal. This invention also includes oral use of nonabsorbable polyamine polymers and amine-related co-agents, such as chitosan, to covalently bind and sequester potentially toxic carbonyl compds. present in the diet, oral use of known antioxidant co-agents and related nutritional factors and use of the primary agent and co-agents in combination with known medicaments for treatment of these <b>neurol.</b> diseases.</p>				
ST	<b>neurol</b> disease drug				
IT	Gangliosides				
	RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)				
	(GM1; pharmaceutical compns. for treatment of <b>neurol.</b> diseases contg.)				
IT	Nitrates, biological studies				
	RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)				
	(drugs; pharmaceutical compns. for treatment of <b>neurol.</b> diseases contg.)				
IT	Gene, animal				
	RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)				
	(for chromosome 17 Charcot-Marie-Tooth disease; identification of)				
IT	Hernia				
	(hiatalp; pharmaceutical compns. for treatment of)				
IT	<b>Parkinsonism</b>				
	(pharmaceutical compns. for treatment of)				
IT	Antihistaminics				

Antioxidants  
Immunomodulators  
Vasoconstrictors  
Vasodilators  
    (pharmaceutical compns. for treatment of **neurol.** diseases  
    contg.)

IT Alfalfa  
Antiarrhythmics  
Anticonvulsants and Antiepileptics  
Antidepressants  
Anxiolytics  
Cholinergic antagonists  
Lecithins  
Mastic (resin)  
Muscle relaxants  
Parsley  
Soybean meal  
Watercress  
Yeast  
RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
    (pharmaceutical compns. for treatment of **neurol.** diseases  
    contg.)

IT Amino acids, biological studies  
RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
    (selenium-contg.; pharmaceutical compns. for treatment of  
    **neurol.** diseases contg.)

IT **Mental disorder**  
    (**Alzheimer's** disease, pharmaceutical compns. for treatment  
    of)

IT Tranquilizers and **Neuroleptics**  
RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
    (antipsychotics, pharmaceutical compns. for treatment of **neurol**  
    . diseases contg.)

IT Rice  
RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
    (bran, pharmaceutical compns. for treatment of **neurol.**  
    diseases contg.)

IT **Nerve, disease**  
    (**diabetic neuropathy**, pharmaceutical compns. for  
    treatment of)

IT **Nervous system**  
    (disease, pharmaceutical compns. for treatment of)

IT **Nervous system**  
RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
    (disease, amyotrophic lateral sclerosis, pharmaceutical compns. for  
    treatment of)

IT Bladder  
    (disease, incontinence, pharmaceutical compns. for treatment of)

IT Hearing  
    (disorder, tinnitus, pharmaceutical compns. for treatment of)

IT **Neurotransmitter** agonists  
RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
    (dopaminergic, pharmaceutical compns. for treatment of **neurol**  
    . diseases contg.)

IT Proteins, specific or class  
RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
    (mercapto-contg., pharmaceutical compns. for treatment of  
    **neurol.** diseases contg.)

IT Ubiquinones  
RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
    (reduced, pharmaceutical compns. for treatment of **neurol.**  
    diseases contg.)

IT Bran

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (rice, pharmaceutical compns. for treatment of **neurol.**  
 diseases contg.)

IT 9027-22-9, Decarboxylase  
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (inhibitors, peripheral; pharmaceutical compns. for treatment of  
**neurol.** diseases contg.)

IT 9001-08-5, Cholinesterase 9001-66-5, Monoamine oxidase  
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (inhibitors; pharmaceutical compns. for treatment of **neurol.**  
 diseases contg.)

IT 50-81-7, L-Ascorbic acid, biological studies 52-67-5, Penicillamine  
 52-90-4, Cysteine, biological studies 55-63-0, Trinitroglycerin  
 56-40-6, Glycine, biological studies 56-45-1, Serine, biological studies  
 57-11-4, Octadecanoic acid, biological studies 59-02-9,  
 .alpha.-Tocopherol 59-43-8, Vitamin B1, biological studies 59-92-7,  
 Levodopa, biological studies 60-23-1, Cysteamine 62-46-4, Thiocetic  
 acid 63-68-3, Methionine, biological studies 63-75-2, Arecoline  
 68-41-7, D-Cycloserine 70-18-8, Glutathione, biological studies  
 70-51-9, Deferoxamine 72-19-5, L-Threonine, biological studies  
 77-92-9, biological studies 87-33-2, Isosorbide dinitrate 150-13-0,  
 4-Aminobenzoic acid 302-79-4, Retinoic acid 357-70-0, Galanthamine  
 364-62-5, Metoclopramide 456-59-7, Cycloandelate 532-11-6, Sulfarlem  
 557-04-0, Magnesium stearate 616-91-1, N-Acetylcysteine 645-88-5,  
 Aminoxyacetic acid 1406-16-2, Vitamin D 4759-48-2, 13-cis-Retinoic  
 acid 6027-13-0, Homocysteine 7235-40-7, .beta.-Carotene 7631-86-9,  
 Silica, biological studies 7757-93-9, Dicalcium phosphate 7782-49-2,  
 Selenium, biological studies 8059-24-3, Vitamin B6 9000-01-5, Gum  
 arabic 9000-07-1, Carrageenan 9000-28-6, Gum ghatti 9000-30-0, Gum  
 guar 9000-36-6, Gum karaya 9000-40-2, Locust bean gum 9000-47-9, Gum  
 mesquite 9000-65-1, Gum tragacanth 9004-32-4 9004-34-6, Cellulose,  
 biological studies 9004-61-9, Hyaluronic acid **9005-25-8**,  
 Starch, biological studies 9007-28-7, Chondroitin sulfate 9012-76-4,  
 Chitosan 9056-36-4, Keratan sulfate 11000-17-2D, Vasopressin, analogs  
 11103-57-4, Vitamin A 11138-66-2, Xanthan gum 12001-79-5, Vitamin K  
 12629-01-5, Human growth hormone 13345-51-2D, Prostaglandin B1,  
 oligomers 15301-69-6, Flavoxate 16679-58-6, Desmopressin 18481-23-7  
 19750-45-9 23210-56-2, Ifenprodil 23288-49-5, Probuco 24305-27-9,  
 Thyrotropin-releasing factor 31329-57-4, Nafronyl 51012-32-9, Tiapride  
**51481-61-9**, Cimetidine 59937-28-9, Malotilate 60719-82-6,  
 Alaproclate 64224-21-1, Oltipraz 66357-35-5, Ranitidine 73590-58-6,  
 Omeprazole 75060-92-3 75364-47-5 76824-35-6, Famotidine  
 81098-60-4, Cisapride 103878-84-8, Lazabemide 105431-72-9, Linopirdine  
 RL: **THU (Therapeutic use)**; BIOL (Biological study); USES (Uses)  
 (pharmaceutical compns. for treatment of **neurol.** diseases  
 contg.)

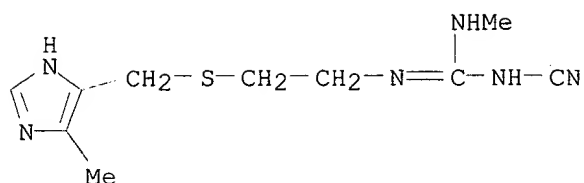
IT 27555-50-6, Poly-N-acetyl-D-glucosamine  
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (.beta.-(1.fwdarw.4), pharmaceutical compns. for treatment of  
**neurol.** diseases contg.)

IT **9005-25-8**, Starch, biological studies **51481-61-9**,  
 Cimetidine  
 RL: **THU (Therapeutic use)**; BIOL (Biological study); USES (Uses)  
 (pharmaceutical compns. for treatment of **neurol.** diseases  
 contg.)

RN 9005-25-8 HCAPLUS  
 CN Starch (8CI, 9CI) (CA INDEX NAME)

\*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

RN 51481-61-9 HCAPLUS  
 CN Guanidine, N-cyano-N'-methyl-N''-[2-[(5-methyl-1H-imidazol-4-yl)methyl]thio]ethyl]- (9CI) (CA INDEX NAME)



L127 ANSWER 28 OF 28 HCAPLUS COPYRIGHT 2003 ACS

AN 1994:672190 HCAPLUS

DN 121:272190

TI Methods for the treatment of bradyphrenia in **Parkinson's** disease using **histamine** antagonists

IN Kaminski, Ram

PA Mount Sinai School of Medicine of the City University of New York, USA

SO U.S., 4 pp. Cont.-in-part of U.S. 5,177,081.

CODEN: USXXAM

DT **Patent**

LA English

IC ICM A61K031-44

ICS A61K031-425; A61K031-415; A61K031-34

NCL 514357000

CC 1-11 (Pharmacology)

FAN.CNT 5

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	US 5352688	A	19941004	US 1992-954258	19920930 <--
	US 5070101	A	19911203	US 1991-655759	19910214 <--
	US 5177081	A	19930105	US 1991-743254	19910809 <--
	US 5453428	A	19950926	US 1993-117503	19930907 <--
	WO 9407490	A1	19940414	WO 1993-US9191	19930927 <--
	W: AU, CA, JP				
	RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				
	EP 665749	A1	19950809	EP 1993-923142	19930927 <--
	EP 665749	B1	20000322		
	R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LI, LU, MC, NL, PT, SE				
	AU 688739	B2	19980319	AU 1993-52929	19930927 <--
	AU 9352929	A1	19940426		
	AT 190840	E	20000415	AT 1993-923142	19930927 <--
	IL 107134	A1	19980222	IL 1993-107134	19930928 <--
	US 5547969	A	19960820	US 1994-314414	19940928 <--
	US 5889033	A	19990330	US 1995-496608	19950629 <--
PRAI	US 1991-655759	A3	19910214	<--	
	US 1991-743254	A2	19910809	<--	
	US 1992-954258	A2	19920930	<--	
	US 1993-117503	A3	19930907	<--	
	WO 1993-US9191	W	19930927	<--	
AB	<b>Neuropsychiatric</b> symptoms of <b>Parkinson's</b> Disease, particularly the symptoms of apathy-amotivation and mental slowing, can be ameliorated by treating a patient suffering from <b>Parkinson's</b> Disease with a <b>histamine</b> H <sub>2</sub> -antagonist (e.g. famotidine, ranitidine) that passes the blood-brain barrier. The H <sub>2</sub> -antagonists may be co-administered with other compds., e.g. <b>histamine</b> H <sub>1</sub> -antagonists or dopamine receptor agonists, which are known to be useful in the treatment of <b>Parkinson's</b> Disease, and can be formulated with such other compds. into a therapeutic compn.				
ST	bradyphrenia <b>Parkinson</b> disease <b>histamine</b> antagonist; famotidine ranitidine bradyphrenia <b>Parkinson</b> disease				
IT	<b>Parkinsonism</b> (treatment of bradyphrenia in <b>Parkinson's</b> disease using				

histamine antagonists)

IT **Blood-brain barrier**  
(treatment of bradyphrenia in **Parkinson's** disease using  
histamine antagonists crossing the blood-brain barrier)

IT Antihistaminics  
(H1, treatment of bradyphrenia in **Parkinson's** disease using  
histamine H2 antagonists and other therapeutic agents)

IT Antihistaminics  
(H2, treatment of bradyphrenia in **Parkinson's** disease using  
histamine antagonists)

IT **Mental disorder**  
(bradyphrenia, treatment of bradyphrenia in **Parkinson's**  
disease using histamine antagonists)

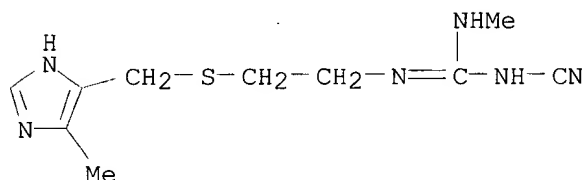
IT **Neurotransmitter agonists**  
(dopaminergic, treatment of bradyphrenia in **Parkinson's**  
disease using histamine antagonists and other therapeutic  
agents)

IT **51481-61-9**, Cimetidine 66357-35-5, Ranitidine 69014-14-8,  
Tiotidine 73147-56-5D, 1,2,5-Oxadiazol-3-amine, derivs. 73590-58-6,  
Omeprazole 76824-35-6, Famotidine 76824-35-6D, Famotidine, homologs  
and isomers 76963-41-2, Nizatidine  
RL: **THU (Therapeutic use)**; BIOL (Biological study); USES (Uses)  
(treatment of bradyphrenia in **Parkinson's** disease using  
histamine antagonists)

IT **51481-61-9**, Cimetidine  
RL: **THU (Therapeutic use)**; BIOL (Biological study); USES (Uses)  
(treatment of bradyphrenia in **Parkinson's** disease using  
histamine antagonists)

RN 51481-61-9 HCAPLUS

CN Guanidine, N-cyano-N'-methyl-N''-[2-[[5-methyl-1H-imidazol-4-yl)methyl]thio]ethyl]- (9CI) (CA INDEX NAME)



=> d his

(FILE 'HOME' ENTERED AT 12:22:18 ON 09 APR 2003)  
SET COST OFF

FILE 'HCAPLUS' ENTERED AT 12:22:34 ON 09 APR 2003

E BAMDAD R/AU

L1 9 S E4,E5  
E BAMDAD C/AU

L2 31 S E3-E7  
E BAMBAD R/AU

L3 2 S E4

L4 1 S E2

L5 32 S L1-L4  
SEL DN AN 15

L6 1 S L5 AND E1-E3  
SEL DN AN L5 11

L7 1 S E4-E6 AND L5

L8 42 S R (1W) ATENOLOL

L9 192 S ALPHA ( ) (METHYLNOREPINEPHRINE OR METHYL NOREPINEPHRINE)  
 L10 243 S ERBSTATIN  
 L11 0 S ICI 11551 HYDROCHLORIDE  
 L12 0 S ICI 11 551 HYDROCHLORIDE  
 L13 0 S ICI 11 551  
 L14 0 S ICI 11551  
 L15 0 S ICI11551  
 L16 0 S INDATRALINE HYDROCHLORIDE  
 L17 32 S INDATRALINE  
 L18 28 S CGS 12066A  
 L19 0 S CGS 12066A DIMALEATE  
 L20 0 S CGS 12066# DIMALEATE  
 L21 22447 S URACIL  
 L22 0 S L21 (L) 5 TRIFLUOROMETHYL 5 6 DIHYDRO  
 L23 0 S 1 ALLYL 3 7 DIMETHYL 8 (1W) SULPHOPHENYL XANTHINE  
 L24 1 S 1 ALLYL 3 7 DIMETHYL 8 (1W) SULFOPHENYL XANTHINE  
 L25 1 S ADSPX  
 L26 1 S 1 ALLYL 3 7 DIMETHYL 8 (1W) SULFOPHENYLYXANTHINE  
 L27 0 S ALPHA BETA METHYLENE ADENOSINE 5 TRIPHOSPHATE DILITHIUM  
 L28 28 S ALPHA BETA METHYLENE ADENOSINE 5 TRIPHOSPHATE  
 L29 4 S HISTAMINE (S) ALPHA METHYL (S) DIHYDROCHLORIDE  
 L30 0 S HISTAMINE 1 METHYL HYDROCHLORIDE  
 L31 3 S 1 3 DIHYDRO 1 2 HYDROXY 5 TRIFLUOROMETHYL PHENYL 5 TRIFLUOROM  
 L32 5 S N6 CYCLOPENTYL 9 METHYLADENINE  
 L33 252 S N6 METHYLADENOSINE  
 L34 34 S S 4 NITROBENZYL 6 THIOINOSINE  
 L35 0 S P P DI ADENOSINE 5 TETRAPHOSPHATE TRIAMMONIUM  
 L36 0 S P P DIADENOSINE 5 TETRAPHOSPHATE TRIAMMONIUM  
 L37 399 S DIADENOSINE(1W)TETRAPHOSPHATE  
 L38 0 S L37 AND TRIAMMONIUM  
 L39 233 S BRL 37344  
 L40 9 S THIOPERAMIDE MALEATE  
 L41 0 S 3 3 4 4 TETRAMETHOXY N METHYL DIPHENETHYLAMINE  
 L42 3 S 3 3 4 4 TETRAMETHOXY(L)DIPHENETHYLAMINE  
 L43 0 S FORMYL(S)HYDROXY(S)PHOSPHONOXY(S)PYRIDIN?(S)BENZENEDISULFONIC  
 L44 0 S FORMYL(S)HYDROXY(S)PHOSPHONOXY(S)PYRIDIN?(S)BENZENE DISULFONI  
 L45 9 S 4 DAMP METHIODIDE  
 L46 14 S FLUOROHEXAHYDRO(S)?DIFENIDOL?  
 L47 77 S P F HHSID  
 L48 53551 S HISTAMINE  
 L49 16 S HISTAMINE (S) METHYL (S) DIHYDROCHLORIDE  
 SEL RN L6

FILE 'REGISTRY' ENTERED AT 13:39:27 ON 09 APR 2003

L50 60 S E7-E66  
 L51 23 S 50-33-9 OR 52-01-7 OR 55-10-7 OR 57-47-6 OR 61-76-7 OR 65-28-  
 L52 18 S 4789-68-8 OR 13153-27-0 OR 13523-86-9 OR 15307-79-6 OR 15676-  
 L53 3 S 147416-96-4 OR 148440-81-7 OR 153587-01-0  
 L54 8 S 147416-96-4 OR 148440-81-7 OR 153587-01-0 OR 109292-91-3 OR 1  
 L55 49 S L51-L54  
 L56 16 S L50 NOT L55  
 L57 3 S 501-75-7/CRN AND CLH  
 L58 1 S 121741-03-5  
 E CGS 12066/CN  
 L59 1 S E4  
 L60 1 S E5  
 L61 14 S L56 NOT L57-L60  
 L62 49 S C6H11N3/MF AND NCNC2/ES  
 L63 7 S L62 AND ALPHA  
 SEL RN 2 4 5 6  
 L64 4 S E1-E4/CRN AND CLH  
 L65 58 S L55,L57-L60,L64  
 L66 13 S L50 NOT L65



L67 71 S L65,L66

FILE 'HCAPLUS' ENTERED AT 14:48:46 ON 09 APR 2003

L68 34870 S L67  
L69 2 S L5 AND L68  
L70 1 S L69 NOT TUMOR MARKER  
L71 2 S L6,L7,L70  
E MINERVA/PA,CS  
L72 725 S MINERVA?/PA,CS  
L73 3 S L72 AND L68  
L74 1 S L71 AND L73  
L75 2 S L71,L74  
L76 24 S L68 AND NEURODEGEN?  
E BRAIN, DISEASE/CT  
L77 105952 S E3+NT  
E E3+ALL  
E E169+ALL  
L78 333107 S E4+NT  
E E52+ALL  
L79 7577 S E4,E3+NT  
E PRION DISEASE/CT  
E E4+ALL  
L80 2209 S E7,E8,E6+NT  
E E16+ALL  
L81 3022 S E7,E6+NT  
E PRION DISEASE/CT  
L82 4 S E11  
E NERVOUS SYSTEM/CT  
L83 275499 S E3+NT  
E E3+ALL  
L84 2 S E4  
E NERVE, DISEASE/CT  
L85 2111 S E49-E53  
E E3+ALL  
L86 27704 S E3+NT  
E E53+ALL  
L87 141419 S E5,E4+NT  
L88 25802 S E25+NT OR E26+NT OR E34+NT  
L89 237038 S E29+NT OR E30+NT OR E31+NT OR E32+NT OR E28+NT  
E ALZHEIMER/CT  
E E9+ALL  
L90 17662 S E6,E5+NT OR E23+NT OR E24+NT OR E25+NT OR E26+NT OR E27+NT OR  
E PARKINSON/CT  
E E6+ALL  
L91 9632 S E4,E3+NT  
L92 3267 S E9+NT OR E10+NT  
E SICKLE CELL/CT  
E E4+ALL  
L93 1818 S E4+NT  
E DIABETES/CT  
E E10+ALL  
L94 52706 S E5,E4+NT  
L95 13963 S E8+NT OR E10+NT  
E AMYLO/CT  
E E41+ALL  
L96 2430 S E2+NT  
L97 67813 S E6+NT  
E DEMENTIA/CT  
E E4+ALL  
L98 3709 S E2  
L99 8544 S L68 AND L77-L98  
E AGGREGAT/CT  
E E6+ALL

L100 7784 S E1  
E AGGREGAT/CT  
E E18+ALL  
L101 4246 S E1  
L102 87941 S E1+NT  
L103 16 S L99 AND L100-L102  
L104 10 S L67 (L) THU/RL AND L76  
L105 64 S L67 (L) THU/RL AND 99  
L106 60 S L103-L105 AND (PY<=2000 OR PRY<=2000 OR AY<=2000)  
L107 53 S L106 AND (PHARMACOL? OR PHARMACEUT?)/SC  
L108 15 S L106 AND (PHARMACOL? OR PHARMACEUT?)/SX  
L109 5 S L106 NOT L107,L108  
L110 55 S L106-L108 NOT L109  
L111 3 S L110 AND 9/SC,SX  
L112 52 S L110 NOT L111  
SEL DN AN 8 27 49  
L113 3 S E1-E9 AND L112  
L114 4 S L75,L113  
L115 5 S L68 AND L100,L101  
L116 8137 S L99 AND (PY<=2000 OR PRY<=2000 OR AY<=2000)  
L117 818 S L116 AND L67 (L) THU/RL  
L118 378 S L117 AND P/DT  
L119 230 S L118 AND US/PC  
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L121 16 S L117 AND (SENILE OR SENILITY OR SENESCEN? OR GELSOIN OR POLYN  
L122 132 S L120,L121  
L123 53 S L122 AND L119  
L124 24 S L123 AND (NEUROLOG? OR NEUROPSYCH? OR ALZHEIMER? OR PARKINSON  
L125 28 S L114,L124  
L126 28 S L125 AND L1-L49,L68-L125  
L127 28 S L126 AND (NEUR? OR NERV? OR ?ALZHEIMER? OR ?PARKINSON? OR COG

2 ANSWER 54 OF 54 REGISTRY COPYRIGHT 2003 ACS

RN 57-47-6 REGISTRY

CN Pyrrolo[2,3-b]indol-5-ol, 1,2,3,3a,8,8a-hexahydro-1,3a,8-trimethyl-, methylcarbamate (ester), (3aS,8aR)- (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN **Physostigmine (8CI)**

CN Pyrrolo[2,3-b]indol-5-ol, 1,2,3,3a,8,8a-hexahydro-1,3a,8-trimethyl-, methylcarbamate (ester), (3aS-cis)-

OTHER NAMES:

CN (-)-Eserine

CN **(-)-Physostigmine**

CN Cogmine

CN Eserine

CN Esromiotin

CN MCV 4484

CN NIH 10421

CN Physostol

FS STEREOSEARCH

DR 511-49-9, 50975-37-6

MF C15 H21 N3 O2

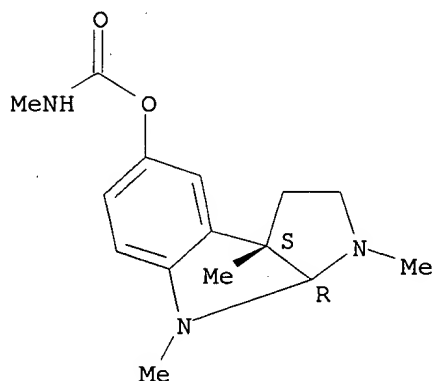
CI COM

LC STN Files: ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, AQUIRE, BEILSTEIN\*, BIOBUSINESS, BIOSIS, BIOTECHNO, CA, CANCERLIT, CAOLD, CAPLUS, CASREACT, CBNB, CEN, CHEMCATS, CHEMLIST, CIN, CSCHEM, CSNB, DDFU, DRUGU, EMBASE, GMELIN\*, HODOC\*, HSDB\*, IFICDB, IFIPAT, IFIUDB, IPA, MEDLINE, MRCK\*, MSDS-OHS, NAPRALERT, NIOSHTIC, PHAR, PIRA, PROMT, RTECS\*, SPECINFO, SYNTHLINE, TOXCENTER, ULIDAT, USAN, USPAT2, USPATFULL, VETU  
(\*File contains numerically searchable property data).

Other Sources: EINECS\*\*, NDSL\*\*, TSCA\*\*

(\*\*Enter CHEMLIST File for up-to-date regulatory information)

Absolute stereochemistry. Rotation (-).



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

3055 REFERENCES IN FILE CA (1962 TO DATE)

33 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

3057 REFERENCES IN FILE CAPLUS (1962 TO DATE)

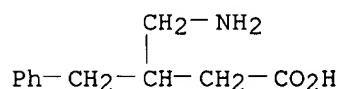
23 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

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ACCESSION NUMBER: 1993:400745 CAPLUS  
DOCUMENT NUMBER: 119:745  
TITLE: Cholinergic sensitivity of irides from donors with various pathological conditions and lens implants  
AUTHOR(S): Patil, Popat N.; Mauger, Thomas F.  
CORPORATE SOURCE: Coll. Pharm., Ohio State Univ., Columbus, OH, 43210, USA  
SOURCE: Naunyn-Schmiedeberg's Archives of Pharmacology (1992), 346(6), 620-8  
CODEN: NSAPCC; ISSN: 0028-1298  
DOCUMENT TYPE: Journal  
LANGUAGE: English

AB In vitro, iris contractions after muscarinic agonists were measured in mg of tension change and the concn. producing 50% of the response was expressed as EC50 .mu.mol/L. Although the av. EC50 value of carbachol in the iris sphincter of the donors with **diabetes** or Parkinson's disease did not change significantly when compared with the control, the max. contraction of the tissue from the diseased state was increased significantly. Thus, in addn. to the well known denervation supersensitivity of the iris-dilator, the iris-sphincter also develops adaptive sensitivity changes. Antimuscarinic drug treatment in some Parkinson's patients interfered with the estn. of supersensitivity in vitro studies. The enhanced response of carbachol at the low temps. or the relative potency of carbachol and pilocarpine in the tissue obtained from the diseased donors was not significantly different from that of controls. Based on EC50 values, the potency of arecoline on the iris was 1/3 that of carbachol. Significantly lower EC50 values of carbachol were found in irides which were in contact with open loop type anterior chamber lens implants compared with those in contact with the closed loop anterior chamber lens implants. Max. responses of irides to carbachol were less when the tissue was in contact with open loop lens compared with those in contact with closed loop anterior chamber implants. Irides from many donors having unilateral or bilateral replacement of the artificial lenses responded with EC50 of carbachol which was approx. equal to that of the contralateral eye. The max. difference between EC50 values of the left and right iris was less than 5 fold. Paired irides with asym. surgical procedures responded unequally to carbachol. The dissocn. const. KB of atropine (1 nmol/L) at 17.degree. was equal to that obsd. at 37.5.degree.. The KB values of himbacine, methoctramine and pirenzepine were 120, 1500, 120 nmol/L, resp. From one tissue to another, there was a spread in the dissocn. const. value of pirenzepine indicating that human iris sphincter may contain a heterogenous population of muscarinic receptors.

L9 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2003 ACS  
 RN 30200-05-6 REGISTRY  
 CN Benzenebutanoic acid, .beta.-(aminomethyl)- (9CI) (CA INDEX NAME)  
 OTHER CA INDEX NAMES:  
 CN Butyric acid, 4-amino-3-benzyl- (8CI)  
 FS 3D CONCORD  
 MF C11 H15 N O2  
 LC STN Files: BEILSTEIN\*, CA, CAPLUS, CHEMCATS  
 (\*File contains numerically searchable property data)



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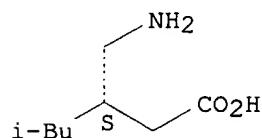
3 REFERENCES IN FILE CA (1962 TO DATE)  
 3 REFERENCES IN FILE CAPLUS (1962 TO DATE)

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 (148553-50-8/RN)

=> d

L10 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2003 ACS  
 RN 148553-50-8 REGISTRY  
 CN Hexanoic acid, 3-(aminomethyl)-5-methyl-, (3S)- (9CI) (CA INDEX NAME)  
 OTHER CA INDEX NAMES:  
 CN Hexanoic acid, 3-(aminomethyl)-5-methyl-, (S)-  
 OTHER NAMES:  
 CN CI 1008  
 CN PD 144723  
 CN Pregabalin  
 FS STEREOSEARCH  
 MF C8 H17 N O2  
 CI COM  
 SR CA  
 LC STN Files: ADISINSIGHT, ADISNEWS, BEILSTEIN\*, BIOBUSINESS, BIOSIS,  
 BIOTECHNO, CA, CAPLUS, CASREACT, CIN, DDFU, DRUGNL, DRUGPAT, DRUGU,  
 DRUGUPDATES, EMBASE, IPA, MRCK\*, PHAR, PROMT, SYNTHLINE, TOXCENTER,  
 USAN, USPAT2, USPATFULL  
 (\*File contains numerically searchable property data)

Absolute stereochemistry. Rotation (+).



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

100 REFERENCES IN FILE CA (1962 TO DATE)

1 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA  
103 REFERENCES IN FILE CAPLUS (1962 TO DATE)

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1 206749-40-8/RN

1 256418-06-1/RN

1 256418-07-2/RN

L11 3 206749-40-8/RN OR 256418-06-1/RN OR 256418-07-2/RN

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L11 ANSWER 1 OF 3 REGISTRY COPYRIGHT 2003 ACS

RN **256418-07-2** REGISTRY

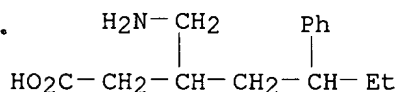
CN Benzenepentanoic acid, .beta.-(aminomethyl)-.delta.-ethyl- (9CI) (CA INDEX NAME)

FS 3D CONCORD

MF C14 H21 N O2

SR CA

LC STN Files: CA, CAPLUS



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

1 REFERENCES IN FILE CA (1962 TO DATE)

1 REFERENCES IN FILE CAPLUS (1962 TO DATE)

L11 ANSWER 2 OF 3 REGISTRY COPYRIGHT 2003 ACS

RN **256418-06-1** REGISTRY

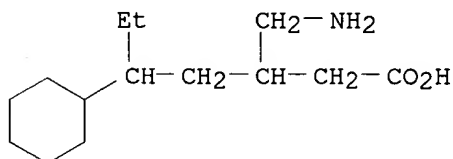
CN Cyclohexanepentanoic acid, .beta.-(aminomethyl)-.delta.-ethyl- (9CI) (CA INDEX NAME)

FS 3D CONCORD

MF C14 H27 N O2

SR CA

LC STN Files: CA, CAPLUS



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

1 REFERENCES IN FILE CA (1962 TO DATE)

1 REFERENCES IN FILE CAPLUS (1962 TO DATE)

L11 ANSWER 3 OF 3 REGISTRY COPYRIGHT 2003 ACS

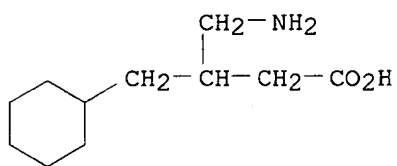
RN **206749-40-8** REGISTRY

CN Cyclohexanebutanoic acid, .beta.-(aminomethyl)- (9CI) (CA INDEX NAME)

FS 3D CONCORD

MF C11 H21 N O2

CI COM  
SR CA  
LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL



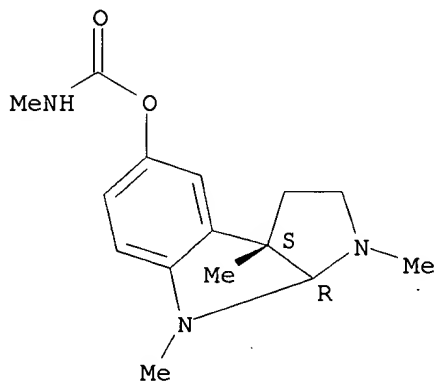
\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

6 REFERENCES IN FILE CA (1962 TO DATE)  
6 REFERENCES IN FILE CAPLUS (1962 TO DATE)

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RN 57-47-6 REGISTRY  
 CN Pyrrolo[2,3-b]indol-5-ol, 1,2,3,3a,8,8a-hexahydro-1,3a,8-trimethyl-,  
 methylcarbamate (ester), (3aS,8aR)- (9CI) (CA INDEX NAME)  
 OTHER CA INDEX NAMES:  
 CN **Physostigmine (8CI)**  
 CN Pyrrolo[2,3-b]indol-5-ol, 1,2,3,3a,8,8a-hexahydro-1,3a,8-trimethyl-,  
 methylcarbamate (ester), (3aS-cis)-  
 OTHER NAMES:  
 CN (-)-Eserine  
 CN **(-)-Physostigmine**  
 CN Cogmine  
 CN Eserine  
 CN Esromiotin  
 CN MCV 4484  
 CN NIH 10421  
 CN Physostol  
 FS STEREOSEARCH  
 DR 511-49-9, 50975-37-6  
 MF C15 H21 N3 O2  
 CI COM  
 LC STN Files: ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, AQUIRE, BEILSTEIN\*,  
 BIOBUSINESS, BIOSIS, BIOTECHNO, CA, CANCERLIT, CAOLD, CAPLUS, CASREACT,  
 CBNB, CEN, CHEMCATS, CHEMLIST, CIN, CSCHEM, CSNB, DDFU, DRUGU, EMBASE,  
 GMELIN\*, HODOC\*, HSDB\*, IFICDB, IFIPAT, IFIUDB, IPA, MEDLINE, MRCK\*,  
 MSDS-OHS, NAPRALERT, NIOSHTIC, PHAR, PIRA, PROMT, RTECS\*, SPECINFO,  
 SYNTHLINE, TOXCENTER, ULIDAT, USAN, USPAT2, USPATFULL, VETU  
 (\*File contains numerically searchable property data)  
 Other Sources: EINECS\*\*, NDSL\*\*, TSCA\*\*  
 (\*\*Enter CHEMLIST File for up-to-date regulatory information)

Absolute stereochemistry. Rotation (-).



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

3055 REFERENCES IN FILE CA (1962 TO DATE)  
 33 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA  
 3056 REFERENCES IN FILE CAPLUS (1962 TO DATE)  
 23 REFERENCES IN FILE CAOLD (PRIOR TO 1967)